

HANDBOOK of MEDICAL TREATMENT

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FOREWORD

This handbook is intended to hold forth the most
 important and practical knowledge of the human
 mind and its functions. It is a guide to the
 study of the mind and its functions. It is a
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WILLIAM T. KERR
 Preface

My dear Sir,
 I am very glad to hear
 of your interest in the

PREFACE TO 6th EDITION

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 June 1953

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Chapter 1

GENERAL ASPECTS OF MEDICAL MANAGEMENT

Selection of all the items for checking during the optimal physical examination of the patient during the following general and particular formulations of the patient's problem

1 Activity status and dependence	Pages 1 and 2
2 Elimination	Page 3
3 Circulation	Page 3
4 Laboratory tests	Page 6
5 Fluid	Pages 5 and 7
6 Symptoms and patient's main	Page 27
7 Diet	Page 46
8 Special measures	See Special Diet

ACTIVITY STATUS

Bed with long bed will be to be a basis for the most method for mobility. It is how to work with it as a danger to it.

The degree of the type of patient should be based on the full condition of the patient's physiological development. The degree of activity should be determined by the patient's ability to perform the daily activities. The degree of activity should be determined by the patient's ability to perform the daily activities. The degree of activity should be determined by the patient's ability to perform the daily activities.

Type of Activity Status

- A Ambulatory For all patients who are able to walk (low)
- B Bed with Bedroom For patients who are unable to walk but are able to move in the bed.
- C Bed with Bedroom For patients who are unable to walk but are able to move in the bed.
- D Complete Bed Rest For patients who are unable to move in the bed.

2 Bed Positions

Undesirable Effects of B & R at

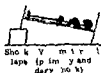
Even in disease states for which bed rest may be employed as a therapeutic measure there are decidedly undesirable consequences which make consideration of early ambulation necessary

- A General. Asthenia muscle atrophy bone atrophy decubitus ulcers vasomotor instability and exsufflation constipation insomnia hypochondriacal trends phlebothrombosis
- B Elderly or Dehilitated Patients. Obstructive pneumonia phlebothrombosis with resultant pulmonary infarction urinary retention from atonic bladder or prostatic obstruction foot and toe deformities (with prolonged rest)
- C Cardiac Patients (Especially those with nocturnal dyspnea) Increased dyspnea when recumbent increased venous stasis and phlebothrombosis
- D Psychiatric Patients. Increased introspection further fixation on physical inadequacies
- E Rheumatoid Arthritis Patients. Deformity ankylosis

Measures to Reduce Hazards of Complete Bed Rest

- A Limitation of narcotic and sedative drugs
- B Deep breathing exercises to prevent orthostatic pneumonia
- C Good nursing care emphasizing frequent changing of position by attendant maintenance of proper elimination and rigorous skin care to prevent decubitus ulcers
- D Improvement of venous circulation by elevation of lower extremities and use of passive bed exercises
- E Foot cradles half bell splints supporting pillows and other measures to prevent physical deformities (see 324) Always have footboard at base of bed

BED POSITIONS AND INDICATIONS



ENVIRONMENT

A suitable physical environment for a given illness must include a consideration of the many personal requirements and idiosyncrasies of the individual patient. The following environmental features should always be considered:

- | | | |
|--------------------|------------------------|---------------|
| 1 Room temperature | 4 Lighting | 7 Elimination |
| 2 Humidity | 5 Social contact | 8 Food and |
| 3 Ventilation | 6 Elimination of waste | 9 Allergens |

CLINICAL OBSERVATIONS

Considerable information of the response as well as diagnostic value may be gleaned from a carefully maintained clinical record. Such information may provide valuable data about the response to treatment, complications and prognosis of the disease and will help in assessing the general comfort of the patient. The following brief clinical data are included for the convenience of the clinician in formulating plans for following the course of the patient. The physician's task is obviously simplified if the patient is hospitalized where adequate nursing facilities are available. Skillful nursing supervision and careful nursing notes of patient's activities, vital signs, medications and other important data make possible more effective management of the patient.

Body Temperature

A Method of Determination and Normal Adult Values

Ar	Arterial Temp	Range of Temp
Rectal	99.6 F (37.5 C)	98.5 - 99.9 F (37 - 37.7 C)
Vaginal	99.6 F (37.5 C)	98.5 - 99.9 F (37 - 37.7 C)
Oral	98.8 F (37.0 C)	96.7 - 99.0 F (36 - 37.4 C)
Axillary	97.6 F (36.5 C)	

B Fever

1 Type

Remittent. Of days or weeks duration with fluctuating periods during which temperature is normal (e.g., brucellosis, tertian malaria). Temperature should be taken tid, qid, or prologed (weekly or monthly) to determine the fluctuating level and average periods.

- b Intermittent. Temperature drops to normal or subnormal level or more in 24 hours (e.g., septicaemia and early tuberculosis). Temperature must be taken qid to determine the variation within the day.

Continuous. Temperature is never normal during 24-hour period (e.g., pneumonia, influenza). Temperature must be taken qid or at times every 2-3 hours to determine its sustained character.

- 2 Cause. Infectious disease, certain drugs, foreign poisons, exposure to external heat, neoplastic disease, disturbance of heat-regulating center and neoplasms.

4 Clinical Observations

C Subnormal Temperatures May be due to profus perspiration hemorrhag shock & creased B M R and mental depression A common cause of recorded subnormal temperature is insufficient time allowed for taking temperature Subnormal temperature may indicate failure in a seriously ill patient and demand immediate supportive intervention In such cases the physician should be notified at once Questionable oral temperatures should be checked by rectal temperatures During afebrile periods it is often advisable to determine the temperatures by the rectal method

Pulse

- A Characteristics which should be noted and recorded include
- 1 Rate (normally 60-80 per minute variable) Compare apical and peripheral pulses in patients with arrhythmias
 - 2 Rhythm Note regularity periodicity and missed beats
 - 3 Intensity Note firmness of the pulse
- B Factor in using the pulse rate include cardiac arrhythmias five cardiac failure hemorrhage shock exertion emotional changes oxygen deficit CO_2 excess increased B M R and certain drugs
- C Factor decreasing the pulse rate include athletic training heart block increased intracranial pressure decreased B M R and jaund

Respiration

- A Characteristic which should be noted and recorded include
- 1 Rate (normal range 14-20 per minute)
 - 2 Volume and depth e.g. deep breathing of Kussmaul type
 - 3 Irregularity and periodicity e.g. alternating hypopnea and apnea of Cheyne-Stokes type
- B Factor in using the rate include error in ease of B M R oxygen deficit CO_2 excess cardiac failure pulmonary disease anemia hemoglobin fever COPD and drugs ketosis emotional states pain nausea and vomiting intestinal necrosis with breathing
- C Factor decreasing the rate include hypnotics and narcotic drugs and increased intracranial pressure

Blood Pressure

- A Details and characteristics to be noted and recorded include
- 1 Position of patient time of recording
 - 2 Systolic level by both palpatory and auscultatory methods
 - 3 Strength and character of pulsations of lower extremities in the presence of brachial hypertension
- B Physiological factors which influence blood pressure include increase in age obesity (if increase is slight to moderate in obese person this is probably due to elasticity of pressure due to abdominal distention excitement rest sitting or standing position (compare with blood pressure in umbilical artery)
- C Pathological factors which influence blood pressure include
- 1 Idiopathic benign and malignant essential hypertension in increased intracranial pressure increased B M R adrenal tumors eclampsia and aortic aneurysm

D Pathological causes of hypotension include hemorrhagic shock, debility as a result of cardiac failure, decreased B.M.R. and hypoadrenocorticism (Addison's disease).

Interrelationship of Vital Signs

- A Pulse Temperature Rectal Temperature
- 1 General rule: Fever (rise) of temperature rise the pulse rate usually rises 10 beats per minute for 98.60/min 99 70/min
 - 2 Diseases in which pulse rate may be low in proportion to fever (relative bradycardia): Typhoid fever, undulant fever, influenza, meningitis, infectious mononucleosis.
 - 3 Diseases in which pulse rate is usually high in proportion to fever (relative tachycardia): Scarlet fever, rheumatism, diphtheria, thyrotoxicosis, subacute bacterial endocarditis, tuberculous terminal or unfavorable pneumonia (pneumonia).

Respiration Temperature Relationships

- 1 General rule: Respiratory rate usually parallels temperature change.
- 2 Exception: In the acute respiratory diseases (relative tachypnea, hyperpnea or dyspnea).

Clinical Blood Pressure Relationships

- 1 General rule: The same factors causing an increase in cardiac rate usually cause an increase in blood pressure.
- 2 Exceptions:
 - a Relative tachycardia: Same as for pathological causes of hypotension.
 - b Relative bradycardia: Renal diseases, benign and malignant hypertension, increased intestinal absorption.

Misleading Observations and Precautions

The following observations are also important in determining the general condition of the patient:

A Fluid Intake and Output: Consideration of the fluid balance of the patient should include the following:

- 1 Clinical utilization of water of hydration.
- 2 Estimation of need for fluids.
- 3 Types of fluid administration.

(For details see Chapter 2, p. 7.)

B Condition of the Skin: Evidence of dehydration (bed or skin) heat, rash, hypohidrosis, dryness, etc.

C Condition of the Mouth, Lips, and Nose: Evidence of dehydration, dryness, etc.

Local condition of the mouth permitting patient to should brush their teeth with a toothbrush daily.

Patient should be given the opportunity to use their mouth after each meal with plain tepid water, physiological saline solution or Alkaline Aromatic Solution N.F. (diluted 2:1).

Caution must be taken that the patient is in a state of hydration.

D Appetite: Question patient regarding appetite, food, drink, etc.

Check to find out if patient is eating, drinking, etc.

Determine reason for rejection of food. A prolonged period of refusal to eat or drink is a sign of a true or relative malnutrition (see text on Diet, p. 46).

6 Laboratory and X ray

- E. Elimination Bed patients are generally prone to constipation. This may be exaggerated by the illness itself, distaste for bland pans diets, and certain drugs. When knowledge of elimination is especially important, gross inspection of all stools passed by the patient may be necessary (see Constipation page 254). Daily inquiry regarding elimination should be made of each patient.
- F. Acceptance or Rejection of Medication Always inquire as to reason for rejection of medication. The patient's objections may constitute valid reason for modification or cessation of drug therapy. Statements as to untoward reactions from medication always deserve careful evaluation.
- G. Sleep and Rest The patient's statements about amount of sleep or rest may vary considerably from known observations. Provide suitable environment for sleeping and resting by insuring a minimum of interruption by professional services, attendants, visitors, and ward mates. Routine sleep inducing drugs should be avoided (see Insomnia page 38).
- H. Mental Reaction of Patient Observe patient's mood and behavior carefully. Watch for mental depression which is often associated with costly, confining, serious or chronic illnesses.

LABORATORY AND X RAY STUDIES

Ordering Laboratory and X ray Studies

The careful and intelligent ordering and performance of laboratory, x ray, diagnostic and other special studies constitutes an essential phase of the management of the patient. The blood count, urinalysis, serological test for syphilis, and perhaps chest x ray should be performed routinely on all hospital patients.

- A. Special diagnostic studies may call for careful planning and integrating with the therapeutic program and must not conflict with the treatment schedule.
- B. Improperly performed or unnecessary laboratory and x ray studies, aside from the discomfort, expense, and inconvenience they cause the patient, may prolong hospitalization.
- C. It must be remembered that certain laboratory studies may require dehydration (e.g., Addison test or pyelograms) when it may be clinically dangerous (e.g., precipitation of renal failure).
- D. Likewise, forced fluids (e.g., P.S.P.) may be contraindicated in the presence of anuria or severe congestive failure, etc.
- E. X ray Plain and dye studies should precede all barium contrast studies, and retrograde barium (nema) studies should precede upper gastrointestinal studies, since reversal of this sequence of studies will cause needless delay.

Chapter 2

FLUID AND ELECTROLYTE THERAPY AND PARENTERAL FEEDING

FLUID BALANCE

In considering fluid therapy it is necessary that the problems of water and electrolyte metabolism be considered independently of each other. The electrolyte is so intimately connected with the maintenance of normal cellular metabolism and basal regulation and with water the maintenance of osmotic pressure in both the extracellular and intracellular fluid compartments. Water in excess of the quantity necessary for maintaining the isotonicity of body fluids is required for normal bodily function. The administration of solution of isotonic electrolytes to the patient cannot be considered as providing available water for the excretion of the same electrolyte. Water must likewise be excreted to keep the solution (urine) almost isotonic.

Daily Obligatory Water Requirements

A considerable amount of water (if from electrolyte) is necessary for normal bodily function. These obligatory water requirements are related to energy expenditure thereby given in the following table.

AVERAGE DAILY WATER NEEDS FOR EXCRETION

Method of Excretion Loss	Volume of Water per 100 Calories of Food	Volume of Water Excreted per Day*
Imperviousible loss (lungs and skin)	44 †	1100 cc †
Sweat (perspirable)	Varies with external temperature. May be high.	0-200 (more at high temperature)
Urine	Varies with amount of waste products to be excreted (see chart page 9)	Average minimum 1000 cc (if perspire more than 1020)
Needs (perspirable) on other than normal		1200-1500
Total (including insensible)		2200-2500+

Based on 2500 Calorie food intake

†For going to clinical estimation the imperviousible loss of water may be calculated as 10 cc /Kg (5 cc /lb) body weight per day

8 Fluid Needs

From the table on the preceding page it is seen that about 1200-1500 cc of water are needed per day in addition to the amount of water necessary for removal of wastes in the urine.

The kidney is the organ which is responsible for regulation of the electrolyte of the extracellular fluid compartment and excretion of the end products of metabolism. In order to preserve the constancy of the extracellular environment the kidney has the ability to concentrate the end products of metabolism. Due to this concentrating ability it is also able to conserve water. This ability to concentrate is limited; the urine can be concentrated to a maximum specific gravity of approximately 1.032. However, the efficiency of the kidney to concentrate falls off rapidly above 1.025 specific gravity. In renal disease states when the ability of the kidney to concentrate is lost, larger amounts of water must be taken or administered in order to remove wastes.

Clinical Evaluation of State of Hydration

- A. History Fluid intake and fluid losses. Sudden weight gain and weight losses should be noted since they most frequently indicate changes in water balance.
- B. Physical Examination Observe body temperature, skin hydration (turgor), tongue, mucous membranes, heart rate and blood pressure.
- C. Laboratory Tests Observe urine volume and concentration, blood counts, hemocrit, serum proteins and N.P.N.
- D. Nursing Notes Weigh patient daily. Detailed intake and output record while patient is under observation are extremely valuable. Output should include not only a record of all measurable fluid losses but also notes of observations on less obvious losses such as increased perspiration and respiration. Keeping these records requires considerable effort so that nursing staff should be notified when this information is no longer needed.

Estimation of Need for Fluid

Because of the necessity of administering fluids to cover the obligatory non-renal excretory mechanisms plus sufficient fluids to permit exertion from metabolic waste by the kidneys, an accurate measurement of the 24-hour urine volume and specific gravity will aid greatly in determining whether sufficient fluid is being administered. Approximately 1.5 to 2.0 Gm. of urinary solids are excreted per day per 100 C of lean body mass. This amounts to approximately 35-50 Gm. of solids per day for the average adult. More concentrated urine suggests generally removal proportionately greater amounts of metabolic wastes, although this relationship does not hold strictly true in diabetes mellitus and certain other diseases which may alter urine concentration by the presence of abnormal amounts of certain substances. The relationships between specific gravity, volume, and urinary solids per 24 hours are shown in the table on the following page.

*2.7 grams of glucose in 3.9 g. mass of albumin will raise the specific gravity of 1000 cc. of urine 0.001 at 15°C.

Sp G Urine	Gm Solids per Liter	Urine Vol /35 Gm Solids (Avg Sp Gr)	Urine Vol /50 Gm Solids (Avg Sp Gr)
1.035 1.030	81.79	400	600
1.030 1.025	79.67	475	685
1.025 1.020	67.55	570	800
1.020 1.015	55.43	715	1000
1.015 1.010	43.31	850	1350
1.010 1.005	31.19	1400	2000

Any sudden and marked change in the present hydration status during the latter portion of the collection period will fail to be reflected in the average of the total 24 hour specimen.

ELECTROLYTE AND ACID-BASE BALANCE

No matter how concentrated both intracellular and extracellular electrolytes in concentration. Although the intracellular and extracellular electrolyte concentrations are approximately the same, most processes in the individual are different. The electrolytes present in the body are grouped into positive ions (cations) and negative ions (anions). The cations are concerned with all the functions of the electrolyte. The anions apparently exert no direct physiological action but are intimately involved with acid-base equilibrium. The following table shows both intracellular and extracellular electrolyte groups as cations and anions.

VALUES OF EXTRACELLULAR AND INTRACELLULAR ELECTROLYTES

Ion	Extracellular		Intracellular	
	mEq/liter		mEq/liter	
	Avg	Range	Avg	Range
Positive (cations)				
Na ⁺	142	135-147	13	30
K ⁺	4	4.6-5.6	40	550
Ca ⁺⁺	5	4.5-5.9	0	0
Mg ⁺⁺	3	1.5-3.0	45	54
Total	155		198	
Negative (anions)				
HCO ₃ ⁻	27	25-30	10	22†
Cl ⁻	103	100-110	3	10
HPO ₄ ⁻	2	1.8-2.3	100	200
SO ₄ ⁻	1		20	96
Org. A	6		0	0
Protein	16		65	
Total	155		198	

The above approximate values are for the

†V. I. M. %

From the previous page it is apparent that the electrolyte patterns of the intracellular and extracellular components are entirely different. Whereas sodium chloride is the main component of extracellular fluid, potassium phosphate possibly as a protein complex is the main component of intracellular fluid. This is of foremost importance in consideration of therapy (see page 17) especially since only the extracellular components are available for clinical measurements.

Since little is known regarding the functions and modes of regulation of the intracellular electrolyte concentration, discussion of the electrolytes must be concerned primarily with the extracellular electrolytes. By a knowledge of the extracellular electrolyte concentration and the mechanisms causing derangement, some inferences can be drawn regarding the intracellular alterations.

VOLUME AND ELECTROLYTE CONTENT OF GASTROINTESTINAL SECRETIONS AND SWEAT

	Avg. 24 hr Vol., ml.	Electrolytes in mEq./L.			
		Na ⁺	K ⁺	Cl ⁻	HCO ₃ ⁻
Extracellular fluid		145	3	111	28
Gastric juice					
Containing acid	2500	10-110	1-32	8-155	0
Achlorhydria		8-120	1-30	100	20
Bile	500	130-160	2-12	90-120	38
Pancreatic juice	700	110-150	2-8	50-95	70-110
Small bowel secretion	100-6000	80-150	2-8	40-135	30
Ileostomy					
Recent	100-4000	100-150	5-30	90-140	30
Adapted	100-500	50	3	20	15-30
Cecostomy	100-3000	50	8	40	15
Feces (loose)	100	<10	<10	<15	<15
Sweat	500-10,000	0-100	0-5	0-100	0

Minor alterations of ion concentration occur in interstitial fluid in response to physical laws governing the production of an ultrafiltrate of plasma.

FUNCTIONS OF THE ELECTROLYTES

The electrolytes of the extracellular fluid serve the principal functions (1) regulation of osmotic pressure and water balance (2) maintenance of ionic equilibrium and (3) idiosyncrasy

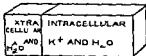
REGULATION OF OSMOTIC PRESSURE AND WATER BALANCE

The osmotic pressure of both the intracellular and extracellular component of the body are at all times equal. In health the osmotic pressure is equivalent to about 310 milliosmole per litre. The total body water is equal to about 40-65% of the body weight with an average of 55% for males and 47% for females. These values are lower in obese and ideal higher in lean individuals. About 15-17% of the body weight is in the extracellular fluid compartment and about 1/3 of this is intravascular. Ament of about the osmotic composition of the extracellular and intracellular fluid is electrolyte difference and this difference is maintained by intracellular enzymic system which remove unwanted electrolyte (eg sodium) rapidly as it diffuses into the cell. Shifts of water between the two compartments is the principal means of maintaining osmotic equilibrium whenever there are alterations in electrolyte or water concentration in the body. The significant alterations that occur counter clinically result in reduced body fluid diagrams. The following diagrams are simplified and fail to show the electrolyte shifts that occur in pathological conditions. In short, electrolyte disturbances complicate the simple equilibrium.

FLUID COMPARTMENTS

Normal

This figure represents total body water with the normal extracellular and intracellular fluid and electrolyte concentrations

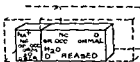


NORMAL

$$\frac{ECF (\text{EXTRACELLULAR FLUID})}{ICF (\text{INTRACELLULAR FLUID})} = K (\text{CONSTANT FOR AN INDIVIDUAL})$$

Simple Dehydration Without Salt Loss

In this condition the electrolyte concentrations are concentrated in the extracellular fluid compartment from the intracellular compartment.



SIMPLE DEHYDRATION

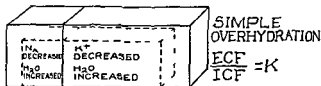
$$\frac{ECF}{ICF} = K$$

12 Functions of Electrolytes

- A Common Clinical Condition Lack of water gastric vomiting
 B Diagnostic Points Marked thirst (probably a symptom of intracellular dehydration) poor tissue turgor high urine specific gravity high hematocrit all extracellular electrolytes may be elevated but proportions normal
 C Treatment Administer fluid without electrolytes Water orally 5-10% glucose in water I.V.

Simple Overhydration

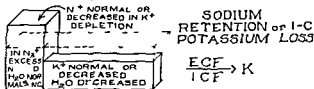
In this condition the electrolytes are diluted the extracellular/intracellular fluid ratio is the same as for normal



- A Common Clinical Condition Excessive fluid intake without salt Excessive electrolyte free fluid administration to patient with oliguria or anuria Excessive posterior pituitary antidiuretic hormone (ADH) production
 B Diagnostic Points Edema low urine specific gravity if patient is in a tinged expect when due to pituitary antidiuretic hormone (ADH) elaboration (in which case low urine output with high specific gravity) low Hct all extracellular electrolytes diluted but proportions normal BP normal or elevated Convulsions if extreme
 C Treatment Usually with holding of fluid and electrolyte

Excessive Sodium Retention or Intracellular K⁺ Loss

This leads to excess fluid in the extracellular compartment with depletion (dehydration) of the intracellular fluid



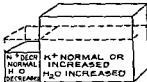
- A Common Clinical Conditions Cardiac failure liver failure with ascites nephrotic syndrome with failure to excrete sodium easily sodium intake administration of metoprolol or ACTH added bilateral disease with inadequate K⁺ intake
 B Diagnostic Points
 1 In Na⁺ retention Edema patient may be thin low hematocrit elevated BP extracellular sodium concentration may be normal or elevated urine specific gravity usually low
 2 In K⁺ depletion Edema may be present serum Na⁺ may be normal but soft dehydrated with severe depletion of intracellular K⁺ Low Na⁺ may be due to intracellular K⁺ transfer or extracellular dehydration (sweat)

C Treatment

- 1 If due to excess Na^+ Retention of dietary sodium or administration of agents to induce diuresis by kidney (digitals, mercurials, acetazolamide (Diamox®), chlorothiazide (Du 1®), etc. Water orally 5-10% gluconate IV (to provide electrolyte free water)
- 2 If due to K^+ depletion Large quantities of K^+ to or chloride into cellular fluid

Exercise 1. Na^+ Loss (Low Sodium Syndrome)

Thirst, diminished extracellular fluid and an increase of intracellular fluid



SODIUM LOSS

$$\frac{\text{ECF}}{\text{ICF}} < K$$

- A Common Cause of Condition: Low sodium intake or use of mercurial diuretics as fluid intake without sodium after prolonged sweating. Addison's disease sodium binding resin therapy sodium loss with gastrointestinal fluid loss
- B Diagnostic Point: Low blood pressure, muscle cramps, low reflexes, mental symptoms of sleep if gravity be affected
- C Treatment: Administration of sodium salts in concentrations greater than 100 mEq/l (e.g. p. 17)

MAINTENANCE OF NORMAL NEUROMUSCULAR FUNCTIONS (Neuromuscular Irritability)

The electrolyte content of the body kept remarkably constant chiefly by the kidney's selective ability to excrete conservatively. The levels and balance of the various positive (+) ions are important in maintaining no malneurosmic irritability and they vary little in health. Wide variations are incompatible with life and symptoms arise when excessive or deficiencies occur.

Relationship of Electrolyte and Nervous Function

The relationship between extracellular ion concentration and neuromuscular function is not clear. Many factors and symptoms indicate a disturbance of the balance of electrolytes which may or may not be detected by the measurement of extracellular electrolytes and fluid ion.

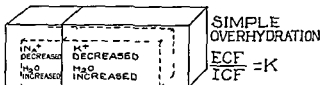
- A Vitamin Effect of Hormones: An example of this is found in the effect of two hormones upon serum potassium levels: namely, dioxycorticosterone and testosterone. Both of the hormones can lower the serum potassium markedly. However, symptoms of potassium deficiency never develop with testosterone. It is believed that this is because the hormone causes an intracellular movement of potassium and that the potassium is removed by the body. Desoxytocorticosterone however

12 Functions of Electrolytes

- A Common Clinical Condition Lack of water gastric vomiting
 B Diagnostic Points Marked thirst (probably a symptom of intracellular dehydration) poor tissue turgor high urine specific gravity high hematocrit all extracellular electrolytes may be elevated but proportions normal
 C Treatment Administer fluid without electrolytes Water orally 5-10% glucose in water I.V.

Simple Overhydration

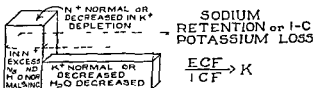
In this condition the electrolytes are diluted the extracellular and intracellular fluid ratio is the same as for normal



- A Common Clinical Conditions Excessive fluid intake without salt Excessive electrolyte free fluid administration to a patient with oliguria or anuria Excessive posterior pituitary antidiuretic hormone (ADH) production
 B Diagnostic Points Edema low urine specific gravity if patient is drinking except when due to pituitary antidiuretic hormone (ADH) elaboration (in which case low urine output with high specific gravity) low Hct all extracellular electrolytes reduced but proportions normal BP normal or elevated Convulsions if extreme
 C Treatment Usually withhold fluid and electrolytes

Example Sodium Retention or Intracellular K⁺ Loss

This leads to excess fluid in the extracellular compartment with depletion (dehydration) of the intracellular fluid



- A Common Clinical Conditions Cardiac failure liver failure with ascites endocrine diseases with failure to excrete sodium excess sodium intake administration of some steroid hormones or ACTH added bilateral gland disease with inadequate K⁺ intake
 B Diagnostic Points

1. Intracellular Dehydration Edema patient may be thirsty when not elevated BP extracellular sodium concentration may be normal or elevated urine specific gravity usually low
2. K⁺ depletion Edema may be present serum Na may be normal but is often decreased with severe depletion of intracellular K⁺ Low Na may be due to intracellular Na⁺ transfer resulting in cellular overhydration (edema)

D Respiratory c dest ind c d by high on nt at ns of CO₂ i
i p ed air leads to c e sed b ght f R and T wa s el va
t n of ST₁ 2 d pr sto of ST₃ and t t m s in e on of
T₁ o T₃ or both

Act of An (Neg t e l s)

The gati on o cerned almost e t ely with osmot c
equ lib i m and pH cha ig s xc pt for the serum i rg e pho
ph rus The pho phorus app rs to b n ded for prope utilis t o
(i ph sphorylat o) of glu o e d pos bly fatty a ds a d so
deq ate qu t t s m y b mp rtant i additio to I V gl co e
whe the l t t i d t d

ACID BASE REGULATION

A d Bas Equ lib ium

The pH of th str cellular fl id during lif is m i ta ed i
7.0 to 7.5 (av ag 7.35 to 7.45). A pH b y nd thi range i in
c mp t bl w th lif The regul t f the pH within s ch narrow
l m ts is th funct n f th buff syst ms f th body p ut up ly
th bic b te buffe At a pH of 7.4 (under no mal cir um
stan s) th s buffe syst m s c mp s d of 1.35 mEq /L of
phy ally diss lv d CO₂ H₂CO₃ d H⁺HCO₃ and 27.0 mEq /L
of B(ba e)HCO₃ whi h g es r tio of 1:20 CO₂ BHCO₃ Th
el tion b p betwe the phy i ally dissolved CO₂ and the H₂CO₃ is
exp s d a f ll w CO + H O \rightleftharpoons H₂CO₃ The on ntr ti
f th d s i ed CO₂ abo i 1000 times s g eat th H₂CO₃
Th H CO₃ d s s t s to H⁺ + HCO₃ to yield the hydrog n ions fo
th buff syst m Si th phys cally d s l d CO₂ is di e tly
r lat d to th partial p e s e of th gas n t ct with th liq id
t b e m l th t the d s ol d CO r te l b lo d nd h c
th hyd g n io o e t at i s d e ctly elat d to i l l CO₂
n ent t (lve la pCO) The int r l t ships an b
r p s t d s s foll w

$$pCO_2 \cong \frac{CO_2}{B + HCO_3} = \frac{H^+ HCO_3}{27.0 \text{ mEq /L}} = \frac{1}{20} = pH 7.4$$

The r t f 1/20 k p th pH i 7.4 rega dl s of th ab olut
quant ti of H₂CO₃ and BHCO₃ pr s t In any probl m inv lving
a disturbanc of s d b s q lib i m a m h bette s lution can
be btain d th efo by knowi g (1) the pH (2) the pCO₂ and (3)
th HCO₃ of the t l b lo d The us l m of obt i ng th
nfo m t n t det mi th pH d tot l CO₂ ont nt of an ero
bi lly il t d and handl d art r l blood Knowing th s two
lve th pCO₂ d HCO₃ an be alc l t d ing th followi g
equ t io s (Tot l CO₂ and HCO₃ n mM/lit pCO₂ in mm Hg)

() To d t min pCO₂

$$pH = 6.10 + \log \frac{(\text{Total CO}_2) - 0.0301 pCO_2}{0.0301 pCO_2}$$

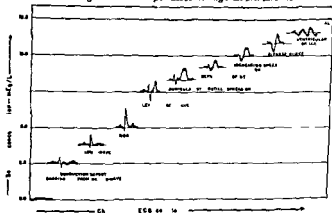
(b) HCO₃ (Tot l CO₂) - 0.0301 pCO₂

marked urinary loss of potassium with intra and extracellular depletion and may lead to symptoms of potassium deficiency

- B Effect of pH on Electrolytes** The pH of the blood is important in determining this intracellular extracellular shift of ions. This is most noticeable in the case of sodium and potassium. In acidosis potassium leaves the cells and tends to be retained by the kidney leading to high extracellular potassium. Sodium may tend to migrate intracellularly during this phase. In alkalosis the reverse occurs potassium moves intracellularly with hydrogen ion while sodium moves extracellularly aggravating the extracellular alkalosis. These changes are shown on p 16

CORRELATION OF THE SERUM POTASSIUM CONCENTRATION AND THE ELECTROCARDIOGRAM

Providing there is no parallel change in Na and Ca



(From K pp Sweet Jawetz and Armstrong Physicists Handbook 10th ed 1958 Lange Medical Publications Los Altos Calif)

Effect of Electrolytes on Electrocardiogram

In addition to the normal range of serum concentration the ECG may be of value in diagnosis of ion abnormalities especially alterations occurring during therapy. Because of the complex interrelationship that occur many times reported a few questionable specificity. The most important are those for Ca^{++} and K^{+} .

- A Ca^{++}** Calcium deficiency prolongs the ST and hence the QT interval. Excess shortens the ST and hence the QT interval. The quantitation of these changes has not been worked out.
- B K^{+}** The potassium changes are best illustrated by the above diagram. There appears to be an interrelationship between Ca^{++} and K^{+} both clinically in terms of symptoms and ECG effects. The symptoms and ECG effects are antagonistic and a deficiency of one may cancel out a deficiency of the other.
- C Mg^{++} and Na^{+}** No specific ECG changes have been described for magnesium or sodium ion change, however the effects of sodium appear to antagonize those of potassium.

Administer half the amount in the first 12 hours; the remainder in the next 24 hours. Give a salt by mouth or by parenteral route.

B. In the case of the replacement of the extracellular fluid, initial replacement can be made especially when parenteral administration is contemplated by estimation of the deficiency in extracellular fluid. This type of calculation obviously in or out of the case of K^+ in this is largely intracellular and deficit may be very large but the amount calculated can be given quite rapidly without dangerous intracellular concentration being produced. Special precautions are necessary in adults with renal insufficiency. Rapid administration may result in hypokalemia. In the case of HCO_3^- replacement it also fails to take into account the intracellular HCO_3^- . For calculation this formula assumes an extracellular fluid content equal to $\frac{1}{3}$ of the body weight. This estimate is not exact, especially in case of disturbed fluid and electrolyte balance.

$$\text{Amount of ion (in mEq)} = \frac{\text{Patient's wt (in Kg)}}{3} \times \left[\text{Normal value of ion (in mEq/L)} - \text{Patient's value of ion (in mEq/L)} \right]$$

Example 100 Kg man has serum CO_2 content of 15 mEq/L. How much sodium lactate is needed to bring serum CO_2 to normal?

$$\text{mEq of } CO_2 = \frac{100}{3} \times (27 - 15) = 20 \times 12 = 240 \text{ mEq}$$

Therefore 240 mEq of sodium lactate is needed by the patient.

Milliequivalent Conversion Factors
for Conversion of Blood Chemistry Findings

Total serum mEq/L of	Divide mg %	val % by
Calcium	2.0	
Chloride (from Cl)	3.5	
(from NaCl)	5.85	
CO_2 Combining Power	44	
Magnesium	1.2	
Phosphorus mM (millimoles)	3.1	
Potassium	3.9	
Sodium	6	

IV FLUIDS AVAILABLE FOR ELECTROLYTE

Many fluids are commercially available and are commonly obtainable. List of the solutions available based on isotonicity and influence on the acid-base balance. Concentrations to be added to each of the electrolyte solutions are used. First half is isotonic and preferably hypotonic when being given intravenously. With solutions of electrolyte in infusion and dilutions.

22 Intravenous Fluids

Neutral Solutions Containing Potassium

See page 25 for precautions with potassium administration
Uses For potassium replacement

- A Dilute Potassium Solutions (Given in 5% glucose solution)
 Two different concentrations are given below

Constituent	% Sol	Gm per liter	mEq / liter of	
			K ⁺	Cl
KCl	0.186	1.86	25	25
KCl	0.3	3.0	40	40

- B Concentrates (To be added to half isotonic or more concentrated solutions only) *Never administer directly from ampule*

1. Concentrate KCl 20 mEq each of K⁺ and Cl in 10 cc of solution (1.5 Gm KCl/10 cc)
2. Concentrate potassium phosphate The addition of potassium phosphate to glucose solutions has been suggested to enhance carbohydrate utilization when indicated (e.g. diabetic acidosis). This may be best prepared by adding the concentrate to 5% glucose solution the pH of the mixture given is suitable for I.V. use. 20 cc ampules contain 60 mEq K⁺ and 32 mM PO₄ per 10 cc
3. Potassium acetate (Quadrate®) alkalinizing but used only for K⁺ effect. 4 mEq/cc in 30 cc ampules

Neutral Solutions of Mixed Electrolytes

Uses To supply other cations besides Na⁺ and K⁺. Their place in clinical practice is still not clearly defined but since losses of electrolytes are rarely single combination replacement may be warranted

- A Ringer's Lactation U.S.P. Compound Injection of Sodium Chloride B.P. (isotonic)

Constituents	% Sol	Gm per liter	mEq / liter of			
			Na ⁺	K	Ca ⁺⁺	Cl
NaCl	0.86	8.6	145			145
KCl	0.03	0.3		4		4
CaCl ₂	0.033	0.33			6	6
Total			145	4	6	155

- B Concentrate of Potassium Magnesium Calcium (KMC®) (To be added to half isotonic or more concentrated fluids) Contains 25 mEq K⁺ 10 mEq Ca⁺⁺ 10 mEq Mg⁺⁺ 45 mEq Cl / 10 cc ampul or

KCl 1.8 Gm }
 MgCl₂ 4.8 Gm } in 10
 CaCl₂ 5.5 Gm }

Solutions of Mixed Electrolyte Yielding Free B

Uses Used where base and potassium replacement is required simultaneously

A Low P l s m L w F B e L c t t d Ring s S i t i n
 U S P Compound Inj c t i o n o f S d m L a t t e B P
 (H r t m a n n s) (i t n c)

Constitu t	% S l	Gm p l t	mEq /l i t o f o n			
			N ⁺	K ⁺	C ⁺	Cl
N Cl	0.6	6.0	102			102
KCl	0.03	0.3		4		4
C Cl	0.02	0.2			3.5	3.5
Sod l t t	0.3	3.0	27			
Total			129	4	3.5	109.5
F r e N ⁺			27			

B High Pota m High F B Lactat d P t s S l n
 I p e c t i U S P (D o w s e l u t i) (VNL) (i s t o n i c)

Co n t i t u t	% S l	Gm p l t	mEq /l i t r f i			
			N ⁺	K ⁺	Cl	La t a t
N Cl	0.4	4.0	70		70	
N l t t	0.6	6.0	53			53
KCl	0.27	2.7		35	35	
Tot l			123	35	105	53
F e N			53			

C V i l n t m d i a t s o l u t i o n c o n t a i n g l e s K⁺ a n d l f e
 b a r i s a v a i l a b l i g G t E l c t o l y t e S l i o n
 W i t h 10% D e t s (B x t e r) i s o t o n i c

Co n t i t e n t	% S o l	Gm p l t	mEq /l i t f o n			
			N ⁺	K ⁺	Cl	Lactate*
N Cl	0.51	5.1	88		88	
KCl	0.03	.9		12	12	
Sod l t t	0.56	5.6	50			50
D x l s	(10)	(100)				
Total			138	12	100	50
F e N ⁺			50			

S l i o i M i d E l t l y t s Y i l d g F A i d i G a t i c E l
 t r o l y t e S o l u t W i t h 10% D t o (B t e) i o t i
 U T r p l e s s i C l l s p l u s f q n t m a l l r
 l o s s e o f N a⁺ a n d K⁺

Co n t i n t	% S l	Gm p l t	mEq /l i t f o n s			
			Na ⁺	P ⁺	NH ₄ ⁺	Cl
N Cl	0.37	3.7	63			63
KCl	0.13	1.3		17		17
NH ₄ Cl	0.37	3.7			70	70
D t	(10)	(100)				
T t l			63	17	70	150
F Cl						70

Sod u m i t t s q l i t t o f e b
 *Ammonium chloride plus f q n t m a l l r
 l o s s e o f N a⁺ a n d K⁺

24 Hypodermoclysis

So called Balanced Electrolyte Solutions

Recently several manufacturers have been distributing so called balanced electrolyte solutions for maintenance therapy. They closely simulate the electrolyte content of plasma except for slightly higher potassium. Their value is questionable however because body losses of electrolytes do not follow the pattern of serum concentrations. In the presence of good renal function it is doubtful that the added base or electrolytes are essential. They have no value as corrective agents and may be harmful in some disease states (e.g. metabolic alkalosis).

ROUTES OF ADMINISTRATION FOR ELECTROLYTES

I.V. Administration

It is most important to exercise great care in administration of I.V. solutions since these alter extracellular fluid concentrations rapidly. This is especially true when giving K^+ and Mg^{++} salts which are exceedingly toxic.

In the presence of normal renal and respiratory mechanisms when swallowing is possible the oral administration of salts is preferred.

Oral Administration

When oral salts are administered usually the negative ions are not too important. See page 20 for the commonly used salts.

PARENTERAL FEEDING AND FLUIDS

When oral feeding is impractical the parenteral routes are the methods of choice.

HYPODERMOCLYSIS

Hypodermoclysis is the subcutaneous administration of large quantities of isotonic solutions usually physiological saline or dextrose in water or saline. Administer at rate of 250-500 ($\frac{1}{2}$ - 1 pt) per hour by use of 1 needle or 500-1000 ($\frac{1}{2}$ - 1 qt) per hour with 2 needles. Usual practical maximum is 3000-4000 cc (3-4 qt) per day. Care must be taken to avoid overdiluting the tissues. This can cause avascularity which may lead to tissue necrosis. When giving glucose solutions $2\frac{1}{2}\%$ in half normal saline is preferred. 5% may be given in water but not to debilitated elderly patients. *Never give glucose in concentrations over 5% to any patient.*

To facilitate absorption hyaluronidase may be added. 250 viscosity units are used per 500-1000 cc of fluid. The mixture may be injected into the tubing of the hypodermoclysis set into the site of insertion of the needle so may be dissolved in the solution. The rate of fluid absorption is increased up to about 12 times.

VENOCLYSIS

The intravenous route of hypotensive administration may be used for fluid and nutrients. Fluid electrolyte carbohydrate protein and vitamin can be administered by this method. The following are the main requirements:

Water

The only way water can be given I.V. with electrolyte to supply the obligatory need for the body is as solvent of 5% or 10% glucose and/or electrolyte in distilled water. NEVER administer plain distilled water I.V. Potassium hydroxides may be in significant quantities of electrolytes (NaCl only) and the end product is

Electrolyte (Minerals)

A Sodium Sodium chloride (NaCl) is the most important electrolyte. Average daily intake 3 to 5 Gm (45-75 gr). Average daily requirement 500-1000 cc (1 pt-1 qt) physiological (0.85-0.90%) saline. If conditions leading to excessive NaCl loss persist added salt may be given. Others do not give over 1 lit (1 qt) physiological saline (8.5-9.0 Gm) daily.

B Potassium If persistent hypokalemia continued for over 3 to 5 days one should polymorph complete electrolyte replacement perily potassium chloride. Average daily body potassium requirement is about 2 to 4 Gm of KCl (25-50 mEq of potassium). This may be achieved by use of potassium chloride potassium by adding potassium chloride to albumin or glucose (page 2). Potassium solutions must be administered slowly (25 mEq/liter of K^+ in 2-3 hours). Never administer potassium I.V. in the presence of poor renal function.

Glucose and Other Sugar

May give 5% or 10% solution. Preferably administered in distilled water but may be given in saline. Never give more rapidly than 0.8 Gm per Kg (1 dr/10 lb) per hour thus the maximum rate of infusion. When given more rapidly than the glycosuria and osmotic fluid loss usually suit. In severe hypokalemia and fluid retention is necessary. Some patients with glucose (20-50%) may be given I.V. very slowly.

Equally potassium and fructose (in water) have been given but have been found to be less effective than glucose and can be administered more rapidly. They differ in that however not marketed. Fructose alone has been used especially in diabetic patients.

Potassium

A Ammonium Usually given Potassium Hydroxide (KOH) (vials) NND and usually administered as 5-6% solution (usually in 5% glucose in distilled water). High concentration of potassium may be added both to this and to the other. This is the only way to administer potassium slowly (about 1 lit (1 qt) in 2 hours). Must be administered slowly. Electrolytes (perily N^+) but have been used should be used if this is important.

Amino acids should not be given when protein itself would be contraindicated (e.g. anuria)

- B Treated Normal Human Plasma U.S.I. has about 7% protein. It is an excellent source of protein. Plasma contains sodium chloride in the same concentration as physiological saline so limit usually is 1 liter (1 qt.) per day. The principal disadvantages of plasma are the danger of producing homologous serum jaundice (even if treated by ultraviolet irradiation) the high cost of the material and possible citrate intoxication with large doses.

- C Normal Human Serum Albumin U.S.P. or Salt poor Serum Albumin. An excellent source of protein. 25 Gm. albumin per 100 cc. solution is equivalent in osmotic pressure to 500 cc. of plasma. This is an excellent way to administer protein in a small fluid volume and with low salt intake by giving the salt poor material. Albumin is very expensive.

Fat

Fat emulsions for intravenous use have been the object of investigation and experimentation for some time. Such a preparation would seem to constitute an excellent means of maintaining nutrition if available although its nutritional value as compared with oral ingestion is not entirely clear. The main problem appears to be that no preparation has yet been offered which is consistently free of reactions, usually severe pyrogenic ones. The emulsions tend to break down especially with freezing or extreme temperature changes.

Vitamins

In prolonged parenteral nutrition one of the more complete B preparations plus vitamin C and parenteral K should be administered. V. R. I. M.

General Indications of Intravenous Alimentation

As soon as practical oral feeding should be started. It is usually impossible to administer adequate calories by intravenous means. The following table outlines the general practical daily physiological limit of intravenous alimentation. The principal limiting factor is the fluid intake. The administration of 3000 cc. of 5% protein hydrolysate in 5% glucose solution would give the following amounts of fluid, electrolyte and nutrient material.

Fluid	Mineral (as NaCl)	Glucose	Prot in Hydrolysate	Calories
3000 cc	Up to 6.0 Gm	150 Gm	150 Gm	1200

Each 50 Gm. of prot in hydrolysate may contain up to 4 Gm. of NaCl, although most products now contain less Na^+ .

Chapter 3

GENERAL SYMPTOMATIC TREATMENT

TREATMENT OF CONSTITUTIONAL SYMPTOMS

PYREXIA (Fever) (code No 003)

Measurements specifically directed toward depression of an elevated body temperature per se are usually unnecessary except for high degree prolonged fevers

A Removal of the Specific Cause of the Fever

- 1 Infection and infectious diseases
- 2 Drug or chemicals Many drugs (eg Monamid) are capable of inducing febrile reactions
- 3 Dehydration Provide adequate oral or parenteral fluid
- 4 Impairment of CNS ability regulating center This poses difficult therapeutic problem Provide for optimal oxygenation and hydration if necessary and prevent excessive hypothermia by a rectal measure if indicated (see below)

B Reduction of the Fever by No Specific Measures When the body temperature is greater than 40°C (104°F) particularly if prolonged the following measures may be utilized

- 1 Increase fluid intake By oral or parenteral routes
- 2 Warm alcohol sponges Cooling is due to evaporation
- 3 Warm moist pads baths These cause peripheral vasodilatation
- 4 Cold sponges Provide prompt cooling of kind as payhology is relatively indifferent with heat loss
- 5 Ice bags Provide local comfort eg forehead
- 6 Antipyretic drugs These drugs are quite effective regardless of fever and hence simultaneous analgesic effect They have the disadvantage that they obscure the clinical picture and may cause undesirable effects such as dyspnea, skin eruptions, hematologic changes, and vomiting cardiovascular depression. Such drugs should be used to be employed cautiously in infectious fevers and preferably in the interfebrile (eg typhoid fever) Atyls Methylated (par) sodium Methylate or c to phenol (phenol) 0.3-0.6 Gm (5-10 g) every 4 hours. Phenol is among the most commonly used and probably the last resort procedure for reduction

SHOCK (Circulatory Failure or Collapse) (code No 0x8)

Shock is a complex and as yet incompletely understood clinical syndrome of peripheral circulatory failure. Numerous pathophysiologic mechanisms are involved in the production of shock such as lack of effective blood volume, alterations in diastolic pressure of peripheral vascular tone, and peripheral vasoconstriction.

of blood oxygenation mechanisms and alteration of the physiological characteristics of the blood. Two principal types of shock must be differentiated clinically.

A Primary Shock (Immediate Shock Fainting or Collapse)

This form of shock consists of transitory insufficiency of circulation and follows relatively suddenly after certain etiologic factors notably those of neurogenic or psychogenic origin. Although shock responds quite promptly to simple supportive measures it is important to remember that primary shock may proceed in idiosyncrasy into the more serious secondary shock. Careful observation of the patient and correction of the aggravating factors are essential. Some of the more common causes of primary shock are:

1. Neurogenic and psychogenic factors. Painful stimuli: trauma, fright, unpleasant sights and odors.
2. Asthenia. Due to anemia, acute infections, chronic illness or to prolonged bed rest. This is noted most frequently when patients assume an erect posture.
3. Certain drugs: e.g. nitrates and local anesthetics.

B Secondary Shock (Delayed Prolonged or True Shock). The onset of this form of shock may be gradual and often is insidious. The classical signs of cold, pale or cyanotic skin, sweating, tachycardia (over 100) and arterial hypotension, although valuable, may appear suddenly and very often represent fully developed shock. Unfortunately advanced shock is often refractory to even the most vigorous anti-shock therapy. Early recognition of shock is imperative. The possibility that shock may occur must always be anticipated. In addition to the above mentioned cause of shock (primary) secondary shock may result from loss of blood (internal or external) loss of plasma; to the serous body cavities (peritonitis) or into traumatized (crushed or bruised) tissues; renal, adrenal, renal cortical insufficiency, dehydration or acute overwhelming infection (e.g. septicemia).

Treatment of Shock

A EMERGENCY MEASURES - act rapidly. The rapidity of shock will vary with the pathophysiological changes which have occurred and which are responsible for the state of shock. Restoration and maintenance of an effective blood volume is the primary emergency measure in combating shock. Dehydration, diminished blood volume, hypoxia, massive infection, hypoproteinemia, acidosis, vascular relaxation and other physiological abnormalities may require special treatment.

1. Body position. Place patient in the shock position (see page 3) unless he has a head injury.
2. Maintain adequate airway.
3. Body warmth. Keep the patient comfortably warm. Avoid chilling or excessive externally applied heat. This will further dilate the peripheral vessels.
4. Pain. Control pain (particularly if severe) promptly by the use of appropriate first aid measures and naloxone. Give morphine sulfate 10-30 mg ($\frac{1}{2}$ - $1\frac{1}{2}$ gr) subcutaneous for pain. Remember that subcutaneous absorption is poor in patients in shock. In case of severe pain morphine sulfate 10-15 mg ($\frac{1}{4}$ - $\frac{3}{8}$ gr) I.V. may be used to great effect.

valge Do not give morphine to unconscious patients patients who have head injuries or those with respiratory depression

Avoid overdosage with morphine but take barbiturates and salicylates for sedation and analgesia whenever possible

- 5 Allay apprehension by reassurance and quietness Pentobarbital sodium 0.1 Gm ($1\frac{1}{2}$ gr) orally or 0.13 Gm (2 gr) subcutaneously if necessary may be of value

- 6 Prevent fluid therapy Replace and maintain adequate blood volume Need for this may be obtained by the history vital signs and hematocrit tests The clinical determination of effective blood volume is difficult however and is a subject for consideration There is no single technique

by which to judge the fluid requirements Response to therapy is a valuable index

- a Small glucose solution Give intravenously 500 cc physiological saline 0.5% dextrose solution 200 cc of 5% physiological saline solution (may be given

orally I.V. while making preparations for plasma albumin or whole blood) Plasma serum albumin and whole blood are obtained in increasing blood volume through the colloidal motopressure effect of dextrose or electrolyte solution

- b Plasma or serum albumin Any of the various plasma preparations has been used for plasma or serum albumin may be employed depending upon the availability Plasma infusion is adaptable may be rapidly set up for administration and does not require preliminary blood typing The quantity of plasma to be given depends upon the stage of shock and the response to therapy based upon the clinical and laboratory data

(1) Impending shock Administer 500 cc plasma intravenously and follow closely clinically and with hematocrit studies to determine effectiveness of plasma

(2) Early or advanced shock Administer 500 cc plasma immediately and repeat with 500 cc every half hour up to total of two liters depending upon clinical course and hematocrit findings If shock persists following such therapy the prognosis is very poor Whole blood If plasma is unavailable transfusion of whole blood may be administered as needed

- d Plasma expanders Evidence during the last few years supports the view that the effectiveness of plasma substitute for emergency treatment of shock The average human body weight is high, high output circulation is necessary viscerally The hypovolemia does not require using infusions of plasma

- (1) Dextrose NND (Epanex® Gt.® Pl. v. l. x®) is water soluble biosynthetic polysaccharide 6% in isotonic saline solution I.V. 500 1000 cc at a rate of 20 40 cc/min USE CAUTIOUSLY in patient with arterial disease Anaphylactic reaction have been reported in order to avoid hemolysis of red blood cells at about 85 mm Hg

(2) **Plazmoid®** a special purified and nonantigenic 5% gelatine solution in isotonic saline I V 500-1000 cc to maintain systolic blood pressure at about 85 mm Hg

- 7 **Vasopressor drugs** These agents are most effective in hypotensive shock without associated decrease in blood volume (e.g. spinal anesthesia syncope and overwhaling intoxications) although they have been stated to be of some value in severe shock due to any cause. They should not be used in lieu of more physiological measures or specific treatment of the cause of shock. In many instances serious question may be raised as to whether the blood pressure elevation produced by the vasopressor drugs has a beneficial or detrimental effect upon the underlying physico-pathological disturbance. (For example the actual influence of the altered peripheral resistance on the blood supply to vital organs is incompletely understood.) Dosage levels for the various agents are empirical and must be carefully adjusted according to patient response (blood pressure and pulse).
- Levarterenol Bitartrate U.S.P. (Levophed®)** 4-16 mg ($\frac{1}{16}$ - $\frac{1}{4}$ gr or 4-16 cc of 0.2% solution) in 1 L. of glucose I V. Avoid extravasation (may cause tissue necrosis and gangrene). With concentrations greater than 4 mg/liter constant supervision and the use of an infusing catheter are required.
 - Phenylephrine Hydrochloride U.S.P. (Neo-Synephrine®)** 0.5 mg ($\frac{1}{120}$ gr) I V or 5 mg ($\frac{1}{12}$ gr) I M or by slow I V infusions of 100-150 mg /liter of glucose.
 - Mephentermine N.N.D. (Wyamine®)** 5-20 mg ($\frac{1}{12}$ - $\frac{1}{3}$ gr) at a rate of 1 mg ($\frac{1}{60}$ gr) per minute by continuous I V infusion or 15-20 mg ($\frac{1}{4}$ - $\frac{1}{3}$ gr) I M.
 - Metamizol Bitartrate N.N.D. (Aramine®)** 2-10 mg ($\frac{1}{30}$ - $\frac{1}{6}$ gr) I M or 15-100 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr) in 250-500 cc of 5% dextrose or 0.5-5 mg ($\frac{1}{120}$ - $\frac{1}{12}$ gr) I V.
 - Methamphetamine Hydrochloride U.S.P. (Vasogyl®)** 15 mg ($\frac{1}{4}$ gr) I M or 5 mg ($\frac{1}{12}$ gr) I V or 35-40 mg ($\frac{1}{2}$ - $\frac{1}{3}$ gr) in 250-500 cc of 5% dextrose by slow I V fusion.

B. Specific (Definitive) Measures

- Anoxia (or hypoxia)** Anoxia is probably present in primary or complicating factors in all types of shock. Therefore oxygen may be considered for most patients in shock. In some patients oxygen may be indicated for other reasons (cardiac failure pneumonia, etc.). However the patient in impending shock is prone to give a death mask so that sometimes serves to increase his apprehension.
- Dehydration** Administer 500-1000 cc of physiologic saline or 5% dextrose solution I V or by hypodermoclysis as needed. As soon as patient can swallow give fluids by mouth. Unless there is positive clinical or biochemical evidence of sodium deficiency avoid administration of more than one liter of physiologic fluid on the first day. Subsequent parenteral fluids may be given as dextrose solutions (see pages 7-10).
- Adrenal cortical failure** Adrenocortical steroid therapy has been found to be effective in shock states associated

with ergo m d l emerg cia Although t e tm f is
mo t p cll c lly applied to sho k of Add n ri is it
may also be of p eta lar v lu in c t in ut lls gie
me ge l and o e whelmi g l o i tion G e Hydro
ortisone (f ee alcohol inf ion con t t fo l V use)
100 150 mg d luted in 1000 c 5% glucos or s lin by slow
I V drip Hyd o o tiso e Sod m S ecia te N N D
(Sol Co taf²) 100 mg m y b giv n in 2 cc ter l w t
r isoto l salin I V over a o minute pe iod it lly
w th sub eq e t dos of 50 mg as q ired

4 Ca diaf l re Digt is dother t atm t f d
f ll are i dicat d o ly fo thos pat ts w th p e e at g
r pr s nng vid ce of c rd failure (eep g 18)

Dg talis i of n val e in shock d e to y oth r ca

5 I fe ti n Immed ate me s s should be tak n to comb t
f ction if p se t Ov whelming inf ct o r sp ble
f p duci g e fficient met b l hanges th body ti s s
to p d spo to ho k P ophylect a t b tics are of
doubt f l val e d m y e be h m f l c cept w th
h d of inf ct l e g eat (g xten i b rns)

6 Hemo hage and ma Alth gh plasma is s lly giv
a an me g ncy meas e i hock compl ati gh mo hag
acut n m m t b o r ted by replacem t with whole
blo d to prev t hyp xia Th q tity of whole blood to b
giv n will dep d pon hematoc it t di

C Ev l tio f Em rg y Th py Constant observation
of patient is imperative The pulse esp atio temper
tu (rectal) and bl od p ss e should b evaluated unrm di
at ly d ve y 15 30 mi tes o off e the e fter until the
is def te imp ov ment of th p iph alci c latio

1 Rapid r ove y If vital s gn ret rn rapidly to n mal
k epp tient und ci ob rvati n b t w th h l d f th
anti hock the py Check vital sign ev ry half h r Pe
f rm h m t rit t d if the is any uspic n whatever
that s onda y h ck exat R m mbe that hem on n
trati n ually p de bloodp e r nd puls changes
Aft r limi ati g pot tial o ext ti g shock p od i g f c
t s the p tie t m y b m ged xp t nly t l it is
es n bly c rta n that d ger h pa s d

2 Prolong d recove y If the v t l igns em in ab orm l fo
ven bri f p lod fte init l me s o sh w ev denc
off r the p og ion of p ph l ircul tory fail e in
stl t fu ther vigor s a t hock the py Blo d hem
globin RBC a d hematocrit ho ld b d t mi d imm d
tely for base line nd ho ld b r p t d s oft n a
ss ry t ev l te th re l f th rapy

PAIN (Ge eral Aspects) (code No 518)

Con ept of pain and p n m hanisms e highly o t o e lal
and fo that s on all i ssific ti s of pain m i quite a bitra y
F p ti al purp es th r a two p ip l typ of pain pe
fici l and d p P in l h a p ly l caliz d in p ficial stru t
and diff sely o poo ly lo l d in d ep rat ct r s D p p i
may es l t in r f r d p P i m y c s a a ult of

multiple types of stimuli acting upon the various body structures. The relief of pain may be achieved by removal of the stimulus or neutralization of the effects of the stimulus and when these are not feasible by dulling or obliterating the sensation of pain.

Analgesic Drugs

A Salicylates The most commonly used of the various analgesics and frequently employed for self-medication.

1 Actions and indication Antipyretic, analgesic, antirheumatic and uricosuric; useful in relieving myalgias, neuralgias, rheumatic headaches and dysmenorrhea.

2 Preparations, dosage and administration

a Acetylsalicylic Acid U.S.P. B.P. (aspirin or ASA) plain or enteric coated 0.3 Gm (5 gr) tablets. Ordinary dosage is 0.3 to 0.6 Gm (5 to 10 gr) every 4 hours per os. 0.3 Gm (5 gr) every 2 to 3 hours is stated to be more effective and lasts in fewer hours than the plain preparation. The plain preparation may cause gastric distress; this may be avoided by administration of the drug on a full stomach or with $\frac{1}{2}$ to 1 tsp of baking soda or other antacid. The enteric preparation is slower acting but it prevents gastric irritation and is also useful for those patients who might be skeptical of the analgesic value of ordinary aspirin. In certain cases it may be necessary to administer the powdered aspirin rectally in a thin starch paste.

b Sodium Salicylate U.S.P. B.P. plain or enteric coated 0.3 to 0.6 Gm (5 to 10 gr) every 4 hours per os.

c Acetylsalicylic acid compound (a purin compound or APC) a synergistic combination

Rx Acetylsalicylic acid 0.22 gr liq

Phenacetin 0.16 gr liq

Caffeine 0.032 gr ss

Sig 1 to 2 tablets every 3 to 4 hours per os

d Aspirin and codeine preparation (see below)

e Analgesic sedative mixtures

Rx Sodium salicylate 10 to 15 gr ss

Elixir of phenobarbital q.s. ad 120 gr ss

Sig 1 to 2 tsp every 4 hours per os

Rx Phenobarbital sodium 0.032 gr ss

Acetylsalicylic acid 0.32 gr ss

Sig 1 capsule every 4 hours per os

f Methyl Salicylate (Oil of Wintergreen) U.S.P. For external use: a liniment to sore muscles, joints. A 10% preparation in oil or ointment.

3 Untoward reactions Usually mild, consisting of drowsiness and somnolence but large doses may cause tinnitus, deafness, blurring of vision, use of vomiting, diarrhea, diaphoresis, headache, delirium. In the patient taking salicylate may cause uric aciduria and may precipitate

B Acetophenetidin U.S.P. Phenacetin B.P. 0.3 Gm (5 gr)

every 3 to 4 hours may be employed in certain cases of salicylate intolerance; in general, however, this drug is more toxic than other analgesic preparations and prolonged use is not advised. Its principal use is in analgesic combination (e.g., APC).

C Colchicine U.S.P. (see Gout page 321)

Narcotic Drugs

Drug which relieves pain and at the same time produces euphoria, sleep or stupor. Pain relief occurs prior to loss of consciousness and catatonic relief of pain of degree greater than that controllable by analgesics or when pain is of type not susceptible to analgesic drug (e.g. visceral pain). All of the following drugs require narcotic prescriptions. These drugs should always be discontinued as soon as the need for them is past.

A Codeine Phosphate U S P B P *16-60 mg*

1. Actions and actions Pharmacologically similar to morphine but of less intensity. CNS depression in ordinary dosage but CNS stimulation in high dosage. diminishes cough reflex and creases bowel motility (constipating). Preferred over morphine for relief of moderate degrees of pain because it is much less habit forming, is much safer and results in fewer untoward reactions.

2. Preparation, dosage and modes of administration

a. Codeine Phosphate U S P B P 0.008 0.065 Gm

($\frac{1}{8}$ 1 gr) orally or subcut. every 3-4 hours p.r.n.

Ordinarily if 0.065 Gm (1 gr) is in the time for analgesia, a strong narcotic since larger doses of codeine are needed by untoward side reactions.

b. Codeine and acetylsalicylic acid (codeine and aspirin)

$\frac{1}{2}$ Codeine phosphate 0.008 0.065 gr $\frac{1}{8}$ 1

Acetylsalicylic acid 0.306 gr v x

Sg. 1 tablet every 3-4 hours p.r.n.

Codeine and acetylsalicylic acid compound

$\frac{1}{2}$ Codeine phosphate 0.016 0.065 gr $\frac{1}{4}$ 1

Acetylsalicylic acid 0.220 gr 11/16

Phenacetin 0.160 gr 1/2

Caffeine 0.032 gr s

Sg. 1 tablet every 3-4 hours p.r.n.

3. Untoward reactions. Allergic reactions such as urticaria,

pruritus, contact dermatitis and anaphylactoid

reactions may occur. Addiction is much less apt to follow use of this drug than use of morphine.

B Meprobene Hydrochloride U S P P Meprobene Hydrochloride *120 f.c.*

B P (Demerol[®] Dolantin[®]) 0.050 0.100 Gm ($\frac{3}{4}$ 1 $\frac{1}{2}$ gr)

orally or I.M. (not subcut.) very 3-4 hours p.r.n. titrated to

be especially useful for pain associated with smooth muscle spasm (especially spasm) although this is disputed. It may be given to individuals who do not tolerate morphine and is less potent than morphine to cause nausea, vomiting and respiratory depression. Analgesic effect is less than with morphine.

Dilaudid[®] and methadone. Addition tendency is finally equal to

C Methadone Hydrochloride U S P (Arm d[®] Dolophin[®]) *5*

0.005 0.010 Gm ($\frac{1}{12}$ $\frac{1}{6}$ gr) subcut. or I.M. every 3-4 hours

p.r.n. provides analgesia of a level similar to morphine but

is slower acting and has a more cumulative action. If titrated

to be especially useful for relief of chronic pain. A

analgesic tolerance develops more slowly than with morphine.

Untoward reactions to the drug are similar to those due to morphine

and the addiction tendency is about the same. The drug

is not tolerated well orally.

D Morphine Sulfate U S P B P This drug remains the most

reliable for potent relief of general local use.

1. Actions and indication: C. t. a. n. v. o. u. s. y. a. t. m. d. e. p. e. s. i. r. u. l. t. i. n. g. i. n. p. w. f. u. l. a. n. a. l. g. e. s. i. a. s. s. o. c. i. a. t. e. d. w. i. t. h. s. d. a. t. i. e. p. h. o. i. a. a. n. d. h. y. p. n. o. i. s. l. e. c. t. i. v. r. e. s. p. i. r. a. t. o. r. y. n. t. e. d. e. p. r. e. s. s. i. o. n. a. n. d. d. i. l. i. n. g. o. r. a. b. l. i. t. y. i. n. f. l. e. x. i. b. i. l. i. t. y. i. n. e. a. s. i. n. t. r. a. c. r. a. n. i. a. l. p. a. i. n. s. u. r. e. h. a. s. m. a. k. e. d. u. s. e. a. t. i. n. g. a. n. d. m. e. l. l. o. r. a. t. o. r. y. e. f. f. e. c. t. s. i. n. c. e. r. t. i. n. p. a. t. i. e. n. t. s. h. a. v. e. m. a. k. e. d. c. o. n. t. i. n. u. o. u. s. u. s. e. s. a. p. p. r. o. p. r. i. e. t. y. a. n. d. u. t. e. r. l. i. m. i. t. a. t. i. o. n. s. i. n. u. s. e. f. u. l. i. n. p. r. o. v. i. d. i. n. g. r. e. l. i. e. f. f. r. o. m. a. c. u. t. e. o. r. p. r. o. l. o. n. g. d. u. r. i. n. g. p. a. i. n. e. p. a. r. t. i. c. u. l. a. r. l. y. p. a. i. n. a. r. i. s. i. n. g. f. r. o. m. c. o. n. d. i. t. i. o. n. s. w. h. i. c. h. a. r. e. o. f. l. e. s. s. t. h. a. n. 10-14 d. a. y. d. u. r. a. t. i. o. n. T. h. e. d. r. u. g. m. a. y. b. e. v. a. l. u. a. b. l. e. i. n. t. h. e. t. r. e. a. t. m. e. n. t. o. f. s. e. v. e. r. e. a. s. t. h. m. a. (g. p. u. l. m. o. n. a. r. y. e. d. m. a. c. h. a. c. a. t. h. m. a. f. l. e. f. t. v. e. n. t. r. u. l. a. r. f. a. i. l. u. r. e.). I. t. i. s. c. o. m. m. o. n. l. y. u. s. e. d. a. n. d. v. a. l. u. a. b. l. e. p. o. s. t. o. p. e. r. a. t. i. v. e. d. r. u. g.

2. Contraindications: M. o. r. p. h. i. n. e. s. e. n. s. i. t. i. v. i. t. y. b. r. o. n. c. h. i. t. i. s. m. s. u. r. g. i. c. a. l. b. d. o. m. i. d. e. (u. n. d. e. r. g. n. o. s. e. d). h. y. p. o. t. h. y. r. o. i. d. e. m. m. o. r. p. h. i. n. e. m. h. e. a. d. i. n. j. u. r. y. A. d. d. i. t. i. o. n. a. l. d. i. s. e. a. s. e. a. n. d. c. i. r. c. u. m. s. t. a. n. c. e. s. w. h. e. n. v. o. r. t. i. c. e. m. y. b. e. d. a. n. g. e. r. o. u. s.

3. Preparations: d. o. s. e. s. a. n. d. m. o. d. e. s. o. f. a. d. m. i. n. i. s. t. r. a. t. i. o. n.
a. Morphine S. l. f. a. t. U. S. P. B. P. 0.008-0.015 Gm (1/8-1/4 gr.) l. i. q. u. i. d. s. u. b. u. t. i. n. a. s. a. s. o. f. s. e. v. e. r. e. a. g. o. n. i. z. i. n. g. p. a. i. n. e. p. a. r. t. i. c. u. l. a. r. l. y. p. a. i. n. a. s. i. t. d. w. i. t. h. i. m. p. e. d. i. n. g. n. e. u. r. o. g. e. n. i. c. s. h. o. c. k. (g. a. s. t. r. o. p. a. n. r. a. t. i. t. i. s). T. h. e. d. r. u. g. m. a. y. b. e. g. i. v. e. n. s. l. o. w. l. y. i. n. 5 c. c. p. h. y. s. i. o. l. o. g. i. c. i. a. l. i. n. i. v. e. n. t. i. o. n.

b. Belladonna l. k. a. l. o. i. d. s. h. a. s. t. r. o. p. i. n. a. n. d. s. c. o. p. l. a. m. i. n. i. n. d. o. s. e. s. o. f. 0.0003-0.0006 Gm (1/200-1/100 gr.) u. b. c. u. t. W. h. e. n. a. d. m. i. n. i. s. t. r. a. t. e. d. i. m. m. e. d. i. a. t. e. l. y. w. i. t. h. m. o. r. p. h. i. n. e. m. y. r. e. d. c. o. l. o. r. a. t. i. o. n. o. f. t. h. e. u. r. i. n. a. r. y. e. f. f. e. c. t. o. f. m. o. r. p. h. i. n. e.

4. Untoward actions: H. y. p. n. o. i. s. (m. a. y. b. e. u. n. d. e. r. t. r. a. b. l. e). p. i. r. a. t. o. r. y. d. e. p. r. e. s. s. i. o. n. n. u. s. a. n. d. v. o. i. d. i. t. i. n. g. s. v. e. r. e. s. t. i. p. t. i. o. n. a. l. g. i. t. a. t. i. o. n. s. p. o. n. s. (u. r. t. i. a. r. i. a. p. r. u. r. i. t. i. s. a. n. d. a. n. a. p. h. y. l. a. c. t. i. o. n. s). A. d. d. i. t. i. o. n. a. l. n. e. c. e. s. s. a. r. y. s. y. m. p. t. o. m. s.

E. D. hydrom. phin. Hyd. o. h. l. o. d. U. S. P. (Dil. u. d. d. s.) 0.005 Gm (1/40 gr.) l. i. q. u. i. d. s. u. b. u. t. i. n. a. s. a. s. o. f. s. e. v. e. r. e. a. g. o. n. i. z. i. n. g. p. a. i. n. e. p. a. r. t. i. c. u. l. a. r. l. y. p. a. i. n. a. s. i. t. d. w. i. t. h. i. m. p. e. d. i. n. g. n. e. u. r. o. g. e. n. i. c. s. h. o. c. k. (g. a. s. t. r. o. p. a. n. r. a. t. i. t. i. s). T. h. e. d. r. u. g. m. a. y. b. e. g. i. v. e. n. s. l. o. w. l. y. i. n. 5 c. c. p. h. y. s. i. o. l. o. g. i. c. i. a. l. i. n. i. v. e. n. t. i. o. n.

Loc. 1 A. th. t. i. s.

A. Cocain. Hyd. o. h. l. o. d. U. S. P. B. P. 10-20% s. l. u. t. i. o. n. f. o. r. t. o. p. i. c. a. t. i. o. n. a. n. d. t. r. a. n. s. p. l. a. n. t. a. t. i. o. n. s.

B. D. b. a. i. Hyd. o. h. l. o. d. N. N. D. C. i. n. h. o. a. f. i. n. Hyd. o. h. l. o. d. i. d. e. B. P. (N. p. a. i. n. e. s. e. n. s. i. t. i. v. i. t. y). I. n. t. e. n. s. i. t. y. o. f. 1.2-2.0% i. n. j. e. c. t. i. o. n. s. l. o. c. a. l. l. y. f. o. r. n. e. u. r. a. l. a. n. a. l. g. e. s. i. a. (N. o. t. o. v. e. r. 100%).

C. Ethyl. Aminob. n. e. t. U. S. P. B. o. i. B. P. 1-20% o. i. n. t. m. e. n. t. a. p. p. l. i. e. d. l. o. c. a. l. l. y.

D. Ethyl. Chl. ride. U. S. P. B. P. s. p. e. c. i. a. l. l. y. u. n. t. i. l. f. o. r. m. a. t. i. o. n. o. f. t. r. a. n. s. p. l. a. n. t. a. t. i. o. n. s. f. o. r. o. s. t. e. o. c. u. r. s.

✓ E. Pr. m. Hyd. o. h. l. o. d. U. S. P. B. P. 0.5-1.0-1.5% o. i. n. t. i. o. n. f. o. r. i. n. f. u. s. i. o. n. a. n. d. t. r. a. n. s. p. l. a. n. t. a. t. i. o. n. s.

Gen. 1 A. th. t. i. s.

Eth. a. d. m. i. n. i. s. t. r. a. t. e. d. b. y. d. e. e. p. i. n. h. a. l. a. t. i. o. n. m. e. t. h. o. d. o. r. a. d. m. i. n. i. s. t. r. a. t. e. d. c. t. a. l. l. y. i. n. g. a. s. t. e. r. i. l. e. o. i. l. 30-90 c. (1-3 oz.).

A. t. i. d. e. a. n. d. A. t. i. p. m. o. d. i. c. s. S. e. p. a. g. e. s. 264 a. n. d. 265.

V. m. o. t. o. D. g. S. e. p. a. g. e. s. 34 a. n. d. 211.

RELATIONSHIP OF AUTONOMIC DRUGS BY MODE AND SITE OF ACTION

These pharmacological relationships are summarized in the following table, illustrating the relationship between the site of action and the mode of action.

AT PARASYMPATHETIC EFFECTORS		AT GANGLIA (P) Symp th t		AT SYMPATHETIC EFFECTORS	
holin E		n olin (m l l d t an t i f t)		Ep phr (d n a l)	
(hist i al int t only)		(hist i al int t only)		Le t (p ph)	
A ty l holi e		A ty l holi e		N i od i e	
M th holine (M h ly ⁶)		M th holine (M h ly ⁶)		Eph d in	
C b h l (D ry ⁶)		C b h l (D ry ⁶)		Synth ti Eph d in lik Am	
B th h l hi id (U e h l i ⁶)		B th h l hi id (U e h l i ⁶)		Ph e ylephrin hydrochl id	
F ri thonium iodide (Furm thid ⁶)		F ri thonium iodide (Furm thid ⁶)		(Ne Syn phrin ⁶)	
h lin i inhib t		h lin i inhib t		Ph ylp opanolamin hyd ochl ida	
Phy oestigmine s li yl t (E l i)		Phy oestigmine s li yl t (E l i)		(P p d line ⁶)	
N oestigmin b mid (oral) (P oestigmin ⁶)		N oestigmin b mid (oral) (P oestigmin ⁶)		P doeph d in	
N oestigmin m thyl ifat (s i) (P oestigmin ⁶)		N oestigmin m thyl ifat (s i) (P oestigmin ⁶)		Amph tamin	
I si roph t (OPP Fl p yit)		I si roph t (OPP Fl p yit)		Amph t m i l f te (B d in ⁶)	
O lam thyley ph ph moid (OMPA)		O lam thyley ph ph moid (OMPA)		D xt mphetamin if t (D d i ⁶)	
Xk p saymop h t Stimulant		Xk p saymop h t Stimulant		Meth mph tamin hyd ochl id (D soxyn ⁶)	
Pill pline		Pill pline		Hyd xamph tami e hyd b omide	
M ari A lin (hd t i al int t only)		M ari A lin (hd t i al int t only)		(P dri ⁶)	
al Bell Jones Alkal da		al Bell Jones Alkal da		C i and P ph (D p ant)	
At opin if t		At opin if t		E gotami e tartir t (Gyn g ⁶)	
B li d n n (lin tur and xt)		B li d n n (lin tur and xt)		E g v n m al t (E got t ⁶)	
S opolamin (Myoscin)		S opolamin (Myoscin)		E got in	
Hyos yamin		Hyos yamin		Dihydroe g tamin (DHE 45)	
2 ymthet All d R l d Dr g		2 ymthet All d R l d Dr g		Dihyd oe g in (DHO 180)	
H mat opin hyd ob omid o al		H mat opin hyd ob omid o al		Synp th lyt Ag t	
H m tropi m thb (N vatri ⁶)		H m tropi m thb (N vatri ⁶)		Dibe amin ⁶	
Amp of opine pho phat (Sy t pa ⁶)		Amp of opine pho phat (Sy t pa ⁶)		Y himbin (hist ic l i t t only)	
E catropi (E pthalimi ⁶)		E catropi (E pthalimi ⁶)		T la lin (P la lin ⁶)	
Adaph nina (T tin ⁶)		Adaph nina (T tin ⁶)		Ph t i m i hydro hl id (R gill ⁶)	
P vat in ⁶		P vat in ⁶		Pipe hydro hl id (B d i ⁶)	
				Hydr i hyd ochl id (Ap u ⁶)	
Dibut lin if t (Dibut ⁶)		Dibut lin if t (Dibut ⁶)			
M thanth li b mid (Banthin ⁶)		M thanth li b mid (Banthin ⁶)			
Oxyph onum bromid (A t yit ⁶)		Oxyph onum bromid (A t yit ⁶)			
Di h m li m thyl if (P i ⁶)		Di h m li m thyl if (P i ⁶)			

- 1 Actions and Indications Central nervous system depression resulting in powerful analgesia as obtained with sedation, euphoria and hypnosis and slight respiratory center depression and dulling or abolition of the cough reflex. It in cases intracranial pressure has marked nausea and emetic effects in certain patients has marked constipating effects. It causes spasm of biliary and uterine smooth muscle is useful in providing relief from acute or prolonged severe pain especially pain arising from conditions which are of less than 10-14 days duration. The drug may be valuable in the treatment of severe cardiac dyspnea (e.g. palpitations, myocardial asthma of left ventricular failure). It is a commonly used and valuable pre-operative drug.
 - 2 Contraindications Morphine sensitivity, bronchial asthma, surgical abdominal distention (diagnosed), hepatic disease, hypothyroidism, morphine poisoning, head injury, Addison's disease and circulatory shock when venous thrombosis may be dangerous.
 - 3 Preparations Dosage and mode of administration
 - a Morphine Sulfate U.S.P. B.P. 0.008-0.015 Gm ($\frac{1}{8}$ to $\frac{1}{4}$ gr) orally as but in cases of ever increasing pain especially pain associated with impending neurogenic shock (e.g. acute pancreatitis) the drug may be given slowly in 5 cc physiological saline I.V.
 - b Bilitinon alkaloide such as atropine and copolamine in doses of 0.0003-0.0006 Gm ($\frac{1}{200}$ to $\frac{1}{100}$ gr) as but when administered simultaneously with morphine may reduce some of the untoward effects of morphine.
 - 4 Untoward reactions Hypnosis (may be undesirable), respiratory depression, nausea and vomiting, gastric contraction, allergic response (urticaria, pruritus and anaphylactoid reactions). Addiction tendency is great.
- E Dihydro-morphine Hydrochloride, U.S.P. (Dilaudid®) 0.0025 Gm ($\frac{1}{24}$ gr) or 0.001-0.004 Gm ($\frac{1}{80}$ to $\frac{1}{16}$ gr) as but every 3-4 hours per oral as indicated to relieve untoward reactions (nausea, vomiting, constipation) than morphine.

Local Anesthetics

- A Cocaine Hydrochloride, U.S.P. B.P. 10-20% solution for topical nose and throat applications.
- B Dibutyl Hydrochloride, N.N.D. C1-hocaine Hydrochloride B.P. (N.P. at 40°C) solution of 1.2-0.00 to 1.1-0.00 injected locally for infiltration anesthesia. (Not over 100 cc.)
- C Ethyl Aminobenzoate, U.S.P. B.N. cocaine B.P. 1-20% ointment for topical use.
- D Ethyl Chloride, U.S.P. B.P. sprayed locally until formation of a thin white frost.
- ✓ E Potassium Hydrochloride, U.S.P. B.P. 0.5-1.0-1.5% solution for infiltration anesthesia.

General Anesthetics

Ether administered by drop inhalation method or administered rectally in vegetable oil B.N. 30-90 cc (1-3 oz.)

Atacide and Antispasmodic See pages 264 and 266

Vasomotor Drugs See page 34 and 211

Autonomic Nervous System Drugs

Disorders of the autonomic nervous system are encountered in a wide variety of diseases both as primary and secondary manifestations. The number and variety of drugs which are used in the treatment of these disorders has recently increased. The pharmacologic relationship of these drugs is shown in the chart on page 34.

PSYCHOTHERAPY

Medical Examination

Although psychiatric diagnoses must be made on psychiatric findings and must not be based solely on exclusion of organic findings, a careful medical examination (history, physical examination and laboratory studies) is of the first significance in approaching a patient with suspected psychogenic disorders. A thorough clinical history can have considerable reassurance value when psychosomatic relationships are discussed with patients. It is really important to point out not only that organic and psychiatric ailments can co-exist but that they are almost invariably interrelated.

- A Avoid either or concept of disease (i.e. either organic or functional) both are usually present.
- B Do not order repeated or prolonged studies merely to impress patient (e.g. extensive investigation of organic trivial).
- C Do not convey unconfirmed suspicion of specific organic disease to patient.
- D Do not hospitalize unless necessary especially if this means exposure to chronically ill patients.
- E Do not neglect medical studies on the preliminary supposition that patient's complaints are functional.

Psychiatric Examination

A careful psychiatric examination usually has both diagnostic and therapeutic value. Expand the usual case history to include inquiry into the patient's motivational difficulties. The thoroughness of this questioning will usually depend upon the given case, but certain basic information is essential. Question slowly and patiently. The examination should include history and observation of the following:

- A Heredity, Family history of psychiatric and constitutional inadequacies.
- B Environmental Factors Development Family childhood training and experiences Family relationships as important friendships social relationships desire and interests as expertises and attitudes (social status and experience person's ambivalence religious attitude).
- C Precipitating Factors Consider especially role of life threatening disease and social problems (financial stress anxiety or hereditary pheasants in way of living death in family overwork and fatigue).
- D Mental Status Of value in distinguishing between functional psychoses and organic complaints. But recognize somatic complaints frequently associated with both neuroses and psychoses.
 - 1 General behavior Appearance speech attention (intensity) Emotion (mood) Anxiety agitation elation or depression.
 - 2 Thought or content Illusions delusions or hallucinations.
 - 3 Sensorium Insight intelligence or orientation memory.

G al Tr t m nt

A T time t f Psy hosomati Sympm ms B fo e dis ov ing
and moving the s of t ns on t may be dvis ble to giv
relief by tre ting symptom All p attive or concrete
medical measures have psychotherapeutic value R pid
re pons to s d ti a d a ti pa mod drug m y be t liz d to
po t o t us pt bility to t tm ta d f ncti n l n tur f d s
s e A c f l explan t f at on l of the th rapy should
foll w ng s mple u de t nd ble te m U ef l m d s
wh h m y mat r ally gme t (but n t repl ce) prog am of
psy h the py in lude t q l z i g o stim l t d g s in
d cated (s e p 4) Ph oba b t a l a d the wer t a
age t hav p d of g at v l in th ma g m nt of p y ho
ge c d u d A widely u ed time t sted a d h g hly s
ce f l p script n s th so lled t t s pa mod m t e
R T ctu of Bell do 10 0 311 s

Elir of Ph b r b t l l a d 120 0 31

Sig f i p t i d 0 m i n a c i 1/2 g l s w t r

B Pl n g of Hyge c L ngr R gtm P ovid for optimal physi
1 d m t i u m t i o l l t r e c o r y
1 I u r d e q u a t e t i t o n w t e g l r b l n e d m e l s
2 P l n a w o r k b l e i v i g h e d u l e w t h a l l w a n e f o r p r o p e
e r c i s r e c r e a t o r a t a d a l p

T t r p t of Sit t i n l Ne osi (Acute emotion al d o d r d e
to u d e r b l x t a l t e s s }

A Permit the patient to tell his troubles (mental illness)
 Let'd go stop as soon as could be time but it is generally
 better to allow the patient to tell his own story

B H l p the p t i t o r c t o I l t situat a l f c t o s
l Util e i g l s o l s i o w e l f a s g i s s d a t d
C t t family o a c i t e s t o o b t d d t i o n l i f o m t o n
d f f i d d c h s e v r o m e t

2 D t d a d a s t a n t o w r d s m p l i f t i o o f p
o l p b l m s C h g e i v o m e t m t a l t i u s
o c u p t i o n l t a t t m y a t t m b e m p s a b l e d
m y c o m p l t a t h t h s m p l i f y p r o b l e m s H l p p t n t
f d h o l u t h t a l l w h o m t o m k h i s o w n d s o

C D t min th p t s a n f o h s t n t h i s t u t
If th p t e t i s s e t e d f g h p b l m o b j e c t l y
l t e d p h i s p h y o c h g i a t t i t u d t o w d t h e s a m p o b
l m m y m a k e h l e a s t i o n m e t o l b l e

D Utilizes blinding (dividing) technique. En g patient to
d lop other ite is pot h bble ad kills pat u
l rly whe patie th exc s of tim fo lf pe p to

E Us ki dly stit de Re u c gg l p su o d
e en dmo to m y b ful as dema ds Av d
p o hng a g w th pati t

T t m t of D e p t d N o s (Chro emotion l d o d s
du to t l confi t lly dat g b k to hldhood }

A R d cat on ie i t te hni sh ld be leg ted to
t an d p y h at ist If s h n t av l bl impl ympto
matic and s pp tv m di al m ssu e d s v the gr t t
con ider tion

B AVOID

- 1 Avoid brutally confronting patient with causal factors of neurotic symptoms
- 2 Avoid premature interpretation of psychiatric data
- 3 Avoid anger toward patient because of failure to improve
- 4 Avoid prolongation of psychiatric study and treatment when it is evident that case progress is unsatisfactory or dangerous. In such cases it is better to refer the patient to individuals more versed in psychiatric methods
 - a Psychoanalytic principles and techniques may be indicated
 - b In inexperienced hands psychiatric meddling may be fraught with dangers. Neuroses are defensive barriers of symptomatic ideas and if these are broken down an emotional crisis may be precipitated or the patient may have undue resentment toward the examiner
- 5 Avoid aggressive psychotherapy during acute or symptomatic phase of patient's disease

Evaluation of the Depressed Patient as a Suicidal Risk

These patients must always be regarded as potential suicides but certain attitudes and responses assist the doctor in determining this possibility

- A Response to Direct Questioning Regarding Suicidal Intent A patient who is afraid he may commit suicide is not likely apt to do so. A patient who feels that he deserves to die or that life holds no hope is apt to commit suicide. These patients may think of suicide but carefully conceal their intentions
- B Doctor-patient Relationship If a patient remains depressed in spite of the doctor's help the patient is a poor suicide risk. If a patient is reassured by the doctor after a reasonable visit he is less apt to be suicidal
- C Reaction to Patient's Normal Environment A patient who has withdrawn from routine living is a poor suicidal risk. The patient who goes on with effort to continue normal daily contacts and work is probably not so apt to be suicidal
- D An increase in neurotic symptom usually indicates the patient is not likely to commit suicide. These are defensive mechanisms

INSOMNIA (Sleeplessness) (cod No 916)

Insomnia is either a failure to fall asleep frequently awakening from sleep or inability to remain asleep for normal periods. Individual sleep requirements however may vary greatly. The causes of insomnia are multiple. Emotional or mental preoccupation is the commonest cause of persistent insomnia.

- A Psychotherapy Direct measure towards correction of deviation of existing anxieties (see page 37). Mutual relationship between and physical therapy method could be included under the heading of palliative psychotherapy

B General Measures

- 1 Promotion of optimal general health
 - a Easily digestible foods in reasonable quantities
 - b Treatment of existing systemic disease
 - c Adequate rest recreation vacations

- 2 R l f of annoying symptom which interfere w th sleep
a Pain (ll types) d Py o i g N l obstruct on
b P itus D b h Dyapnea and orthopnea
c Na a f Cough i Urinary d stu banc
- 3 Quiet pre b dtume activity R t tion of x iting act vit e
specially in th pre bedtim pe iod is an individ al matt r
It i prob bly ad i able for susc pt bl individuals to a oid
exciting or thought p voking reading gam s d am or
movi s for a p iod of l 2 o mo e hours before bedtm
Encour ge light read g and othe non exciting activit s
- 4 Re tri t ion of stimulati g b v ag s and drugs sp c ally
aft 3 00 p m e g te offe tob c o ephedrin lik
d g and amph tamine compounds
- 5 Provision of ad quate sl eping f c l ti g omfortabl
b d and a quiet and da k room with s t bl nt lat tern
p tu and hum dity
- 6 Wa m bath b fore b dtum may h ve a elaxing eff ct
- 7 Warm milk tak n at bedtm m yal o h ve a relaxing effect

C Sed t e a d Hyp ot Agent Th outline e of hyp otic
drug to o t o i m n a i n t o ly mp op b t m y also be
dang rous b use of hab tuati n d i r m d ge of f iling
f m bed te The follow g g is may b u d

- 1 Win (weet h y o sim la) 60 c (2 o) or lly h
- 2 Wh k y 30 cc (1 o) d l ted w th wat o ally h s
- 3 Ph nob rbit l U S P Ph ob hito B P o lly h
slow (30 60 minutes) ct o and a p lo g d eff ct (6 8
ho s) and pati t pt to h e a hango e It e
ct d by kid y and so o t ind ted r l ineuff
cien y

Ph ba b tal U S P Ph nob rbiton B P 0 015
0 03 Gm (1/4 1/2 g) b d q d as a s d i v d g
th dayt m m y d an lty o te s n uffic ally to
obvat th n ty fo hypn t do of b b t ales
t ght D ow d d i e s m y b ov r om by
ca f l ad j stme t of th d g o d i ng to d dual
quir m nts S ll d f il e of th b b tur t a a
t quill and pharm olog d eff ct d t ty
a e freq tly d to improp d sag ad j tm t Th
hyp tic dos f ph obarb t l 0 1 0 2 Gm (1 1/2 3 gr)
o lly h p r

- b Ph nob bit l Elixir U S P 16 3 cc (4 8 d) n
tain 16 mg p r 4 c orally h p n
Ph nobarbit l Sod um U S P Phen ba biton Sodium
B P 0 065 0 1 Gm (1 1/2 gr) 10* s luti
sub ut p n

- 4 P tob hital Sod um U S P P tob rbit o e Sod um
B P orally h m r pid effect (15 30 minut s) and
horter d stion of acti (4 6 hou s) than phenob rbit l
It is ex ted by th liv and i th fo ont andi ated
in h p th lns fici y
a Pentobarb t l Sodium U S P P nt b biton Sodium
B P 0 1 0 2 Gm (1 1/2 3 gr) o lly h s p n
b P tob bit l Sodium Elixir (N C A) 16 32 (4 8 d)
ont in 20 mg p 4 c o lly h s p n
c P t barbit l Sod um r t l pposit y (N C A) 0 13
Gm 1 2 in rt d ect lly h s p n

d Pentobarbital Sodium Sterile U S P 0.5 Gm administered as a freshly prepared 5% solution I M or I V (slowly and not more than 1 cc per minute)

Toxic reactions to barbiturates include excitement and delirium (especially in children and in elderly debilitated or febrile patients) drug addiction barbiturate dermatitis and circulatory and respiratory depression (see p 536)

5 Chloral Hydrate U S P B P (12.5% Sol) 2.4 cc (0.45-0.50 Gm) orally h s p r n

6 Sodium Bromide Elixir N F (17.5% Sol) 1.2 dr h s p r n

7 Paraldehyde U S P B P A useful agent since the clean stock solution is stable and can be used for oral or rectal I M or I V administration as needed. The drug has an unpleasant odor. It may be used in delirium.

a Oral 4-16 cc (1-4 dr) in cracked ice with milk fruit juice or whiskey

b Rectal 16-32 cc (4-8 dr) in 30-60 cc (1-2 oz) of a vegetable oil (1:2 dilution)

c I M 5-10 cc (1-2 1/2 dr) (Preferably deep in buttocks)

d I V 1-2 cc (15-30 min) in triple volume sterile saline very slowly CAUTION May cause respiratory arrest or pulmonary edema

8 Antihistamine The sedative drug is being used widely for its sedative effect (see p 45)

9 Tranquilizing drugs See below and pp 42-43

TRANQUILIZING DRUGS

(See table on pp 42 and 43)

The tranquilizing or ataractic drugs have been accorded a wide acceptance. New preparations are being introduced at a rapid rate. They are being used in the symptomatic treatment of many psychomotor dysfunction disorders for which the barbiturates, barbitals and similar drugs have been successfully employed in the past. There is no doubt that these agents have made an important contribution particularly in the field of psychiatry. If the popularity which has surrounded the introduction of these new agents has done nothing else, it has increased professional and public optimism toward serious psychiatric disorders and brought about a closer relationship of psychiatry with the general medical community. However, a cautious and conservative approach toward the routine widespread use is indicated. The widely enthusiastic report of early results has been somewhat qualified by the results of some recent studies and the important problem of toxicity and long-term effectiveness is a serious concern in the development of all tranquilizers.

The tranquilizing agents have been used with caution cooperatively because of their tendency to aggravate the hypotension which may be encountered in the administration of the anesthetic agents.

None of these drugs can take the place of conventional psychiatric techniques aimed at determining and removing the causes of the illness.

Ph th C mp d

Pharmacologically the compounds most closely fulfill the
 point of tranquillizing drug in that they serve to reduce psych-
 ic without pronounced sedative or hypnotic effect. Up to
 the present year not a word of toxicity principle effects to
 date have been observed with Chlorpromazine U.S.P. (Thorazine®)
 the first of the group to be introduced although it would be likely
 that the more recently marketed related compounds would be
 implicated with the same cut (See table on pages 42-43)

R w lf Prepa ti s

Rawlf's alkaloids are generally recommended for a wide
 variety of conditions. A number of these are now being
 employed on a more limited basis for the treatment of
 various types of mental and emotional disorders. To
 this day the products are fortunately not uncommon
 (See table on page 42-43)

M p ob m t N N D (Miltow® Eq 1)

This drug made it difficult to give a clear how-
 ever although many have found it a useful drug for the treatment of
 anxiety states. The product has demonstrated
 sedative and hypnotic effects. Even though the product
 also has a direct effect on the central nervous system. To
 avoid effects may be recommended the appropriate doses
 of the drug (See table on pages 4-43)

Oth N b b t t T q il

The crynmb of the drug is a definite limitation
 upon which may make any use of it difficult (See
 table on pages 42-43)

ALLERGIC DISORDERS

Allergic disorders may be manifested by any of the
 following: local reactions, any organ system of the body
 The effect may be a local or systemic and may be
 induced by a number of offending agents (antigens). Many
 of these are of all kinds of the human body
 and it is possible all of these. Several different
 varieties of all types of hypersensitivity reactions

N m l Alle gi (C ke)

Dilution test: though not a test with the antigen
 in order to appraise the extent of allergic percentage
 of normal individuals without visible symptoms
 The diagnosis may be readily obtained by appropriate
 skin testing the appropriate (CAUTION)

PSYCHOTROPIC (MOOD ALTERING) DRUGS
TRANQUILIZING (ATARAXIC) DRUGS

Name	Usual Adult Dosage	Indications	Contraindications	Side Effects
AUTONOMIC SUPPRESSANTS				
Phthalimide Compound Chlormethide HCl U.S.P. (Thalimide)	10-25 mg bid qid	Tardipathy dilatation hypertension myocardial infarction renal insufficiency	Contraindications CNS depression tuberculosis hypertension pregnancy (?)	Drowsiness blurred vision undiagnosed hypertension hypertension dry mouth tinnitus hypertension hypertension
Proprietary HCl NND (Spartan)	75-50 mg bid qid			
Meprobamate (Miltrex)	25-50 mg bid qid			
Proprietary NND (Cromol)	5 mg tid qid			
Barbiturate Alk. Salt Risperidone NND (Spartan)	0.1-0.75 mg tid qid			
Risperidone NND (Risperidone)	50-500 mg tid qid			
Risperidone (Modip)	0.25-0.5 mg bid			
Diphenhydramine Compounds A cyclohexanol (Frederic)	20 mg tid			
Hydroxyzine HCl NND (Atax)	25 mg tid qid			
Benzodiazepine NND (Spartan)	1 mg tid			
TRIAL RELAXANTS (Barbiturates)				
Substituted Propanediol Mephobarbital NND (Miltrex Equivalents)	400 mg tid qid			

44 Allergic Disorders

- A Serum sickness
- B Drug anaphylaxis (see below)
- C Dermatitis venenata (see p 87)
- D Tuberculous sensitization (see p 129)

Atopic Disorders (Cocci)

These natural or spontaneous allergic occur about 10% of the population often with a hereditary background. Antigenic etiology is much more obscure than the cause of the normal allergies. Determination of the allergens is much more difficult since complete reliance cannot be placed upon clinical history skin tests or elimination diets. Clinical or therapeutic trials are often misleading. Eosinophilia is characteristic but certainly not pathognomonic of atopic disorders.

- A Hay fever (allergic rhinitis) (see p 112)
- B Eczema (see p 70)
- C Urticaria (see p 78)
- D Angioneurotic edema (see p 78)
- E Allergic purpura (see p 244)
- F Allergic migraine (see p 344)
- G Allergic asthma (see p 115)

Anaphylactic Reaction (Anaphylactic Shock)

Anaphylactic reactions are the immediate shock like and frequently fatal reactions which occur within minutes after parenteral administration of foreign sera or drug. Although there is usually also a history of previous exposure to the foreign body, the severe reactions undoubtedly represent increased hypersensitivity. Anaphylactic reactions to sera, penicillin and other antibiotics and practically all other repeatedly administered parenteral therapeutic agents may occur. It is for this reason that potentially life-saving drugs should not be administered indiscriminately by oral, topical or parenteral routes. Likewise it is probably best to have emergency drugs available in the event of an anaphylaxis. Penicillinase injectable (Neutrapen®) 800,000 unit I.V. followed immediately by 800,000 units I.M. has been reported to be of possible value in penicillin anaphylaxis. Experience with this drug is limited.

Symptoms include apprehension, paresthesia, generalized urticaria or edema, hives, cyanosis, wheezing, cough, incontinence, shock, fever, dilatation of pupils, loss of consciousness and convulsions. Death may occur within 5-10 minutes.

A. Emergency Treatment (URGENT)

- 1 Epinephrine Hydrochloride U.S.P. 1 cc of 1:1000 solution (1 mg) I.M. IMMEDIATELY and repeat in 5-10 minutes as directed. It may be necessary to give 0.1 to 0.4 cc of 1:1000 solution diluted in 10 cc saline SLOWLY I.V.
- 2 Place in horizontal position. Keep warm.
- 3 Maintain adequate airway.
- 4 Diphenhydramine hydrochloride (Benadryl®) 50 mg I.V. immediately.
- 5 Positive pressure oxygen therapy (see p 148).
- 6 Amlopylline 0.24-0.48 Gm I.V. 10-20 cc of saline SLOWLY I.V. may be of aid.

B. Prevention

- 1 Precautions. Be aware of the danger. Avoid using potentially dangerous gauless than is a definite need. Avoid giving drugs to patient with a history of hay fever.

Chapter 4

DIETETICS AND NUTRITION

A diet must supply the following essential components. These requirements can be normally met by including the basic foodstuffs as outlined on page 47. (The dietary components are altered as indicated for the needs of the individual.)

- A Calories for energy (supplied mainly by CHO intake)
- B Protein for growth, development, tissue repair, and energy
- C CHO for energy and for prevention of ketosis
- D Fat for essential fatty acids and energy
- E Minerals and vitamins for maintenance of optimal tissue function and electrolyte equilibrium
- F Water for absorption, transport of foods and waste products and excretion

Modifying the Basic Diet

Personal eating habits, racial and religious restrictions, expenses, and geographic availability of food must be considered in the preparation of any diet. Otherwise, the basic diet must be modified as follows:

- A Increased All or part of the diet must be increased to compensate with reference to the activity of increased metabolism (as in thyrotoxicosis, tissue injury, and fever). (BMR is increased 13% with each 1°C, 7% with each 1°F of fever.)
- B Decreased In obesity, the diet should contain calories, CHO, and fat in decreased amounts.
- C Restricted Some diseases states require specific restrictions or variations of one or more of the basic dietary constituents (see page 54).

PLANNING AND WRITING A DIET

The planning and writing of a diet can be accomplished by following the steps numbered below.

Prescribing the Diet

	Pag
1 Calculate the caloric need of the patient	47
2 Include the basic foods for a well balanced diet	47
3 Select the type of diet to be employed	48
4 Calculate the variation of the dietary component as specified by the diet	48

Selection of the Dietary Components and General Instructions

5 Determine the potential needs and select foods to be used	49
6 Select the carbohydrate foods for the diet	51
7 Determine the fat requirements and dietary sources	52
8 Determine the need for vitamin supplements	52
9 Determine the need for mineral supplements	53
10 Arrange the number, frequency and time of feeding	53
11 Give detailed instructions to the patient	53

STEP 1 - CALCULATE THE CALORIC NEEDS OF THE PATIENT

The caloric needs vary with age, weight, activity, and nutritional status (see below). The caloric needs must be adjusted also to meet variations caused by disease or disorder (see page 46).

APPROXIMATE DAILY CALORIC ALLOWANCES (N R C 1953) Healthy persons 25 years of age U S A (mean temp 10° C 50° F)

Caloric for Standard	Vary Caloric Allowances as Follow
Men 65 Kg (143 lb) 170 cm (67 in) 3200 Calorie	Weight For each 5 kg over standard add Men + 5% Women + 6% For each 5 Kg under standard subtract Men - 6% Women - 9%
Women 55 Kg (121 lb) 157 cm (62 in) 2300 Calorie	Temperature (Average Environmental) For each 10° warm decrease 5% For each 10° cooler increase 5%
Pregnant 2700 Cal Lactating 3300 Cal	Activity Vary to maintain weight

STEP 2 - INCLUDE THE BASIC FOODS FOR A WELL-BALANCED DIET

The well balanced diet should always include serving of the following food stuffs unless contraindicated. These basic foods must be kept in mind while writing the diet and are conveniently used as the nucleus of formulating most diets. The remainder of the diet is made up of increased quantities of foodstuffs from any of the groups. The high CHO foods supply needed also of energy and are the least expensive.

BASIC FOODSTUFFS

Foodstuff	Basic Requirement	Weight or Volume
High protein Food		
Meat, fish or fowl or beef or dried liver	1 serving	3-4 oz (90-120 Gm)
Eggs	1 or 2	1 1/2 oz (45-90 Gm)
Milk whole	Adult 2 glasses Children 4 gla	16 (480 cc) 32 oz (960 cc)
Whole grain (wheat) cereal	1 serving	1 oz (30 Gm)
Bread	2 slices	2 oz (60 Gm)
Vegetable		
1 root	1 serving	4 oz (120 Gm)
1 whole leafy	1 serving	2 oz (60 Gm)
1 other	1 serving	3 (90 Gm)
Fruit		
1 fresh (citrus tomato)	1 medium sized	3 oz (90 Gm)
1 other	1 serving	3 oz (90 Gm)
Fat		
Butter or tallow margarine	2 tablespoons	1 oz (30 Gm)

Individual weight (a background)

†Weight and limit of food stuff and not to the amounts of CHO of it they will yield

48 Types of Diets

When preparing the food always make the servings attractive to sight taste and smell and serve at the proper temperature. The best planned diet is useless unless eaten by the patient.

STEP 3 - SELECT THE TYPE OF DIET TO BE EMPLOYED

After having calculated the basic caloric needs in Step 1 use the following table to help select the type of diet for the disease in question. Descriptive details of these diets will be found on pages 54 to 59.

THE PRINCIPAL TYPES OF DIETS

Disease or Disorder	Diet
Gastrointestinal Peptic ulcer Functional disorders	Modified Sippy Bland (low residue soft consistency non stimulating)
Gallbladder disease Liver disease Constipation	Low fat and non gases forming High protein high CHO High fiber
Cardiovascular Congestive failure Hypertension	Low sodium (salt) Low sodium (less than 300 mg /d y)
Metabolic Diabetes	Usually high protein with moderate CHO restriction (see page 57)
Obesity Wasting and malnutrition	Low caloric adequate protein High caloric high protein high vitamin
Renal Nephritis	Low but adequate protein 0.5 Gm /Kg (0.25 Gm /lb) body weight per day plus total albumin lost in urine
Allergic Food allergy	Special elimination

These diets vary in the number of calories and/or in the amount of one or more of the dietary components. The next step is to calculate these variations.

STEP 4 - CALCULATE THE VARIATION OF THE DIETARY COMPONENTS AS SPECIFIED BY THE DIET

After determining the basic caloric needs and selecting the type of diet use the following table to calculate the number of calories and the amount of each dietary component for the diet. The remainder of the total calories not supplied by the fixed components of the diet may be made up with unrestricted foods.

VARIATIONS OF DIETARY COMPONENTS

Component	Average Diet	High or Increased	Low or Decreased
Calories (Energy)	Variable (See Step 1 page 4)	25-35% more or less than for maintenance	25-35% less than for maintenance
Protein	1 Gm./Kg. (0.5 Gm./lb.) body wt./day (See Step 5 below)	2-4 Gm./kg. (1-2 Gm./lb.) body wt./day (300 Gm. or about 1 lb. per 1 m.t.)	0.5 Gm./kg. (0.25 Gm./lb.) body wt./day (Should not be below)
CHO	50% of calories as CHO	75% or more of calories as CHO	About 25% of calories as CHO
Fat	About 100 Gm. per day	150-250 Gm. per day	70 Gm. or less per day
Vitamins	Supplied by well balanced diet (See page 47)	In case of high vitamin foods or supplement	Not indicated
Mineral			
Sodium	5-20 Gm./day	Above 20 Gm./day	0.2-2.0 Gm./day
Chlorine	0.1-1.5 Gm./day	Above 3 Gm./day	0.2-0.5 Gm./day
Note: If iron is abundant the potassium may be as low as 0.3 Gm./Kg. (0.15 Gm./lb.) body wt. per day			

Having formulated the dietary prescription (Step 14) prepare the actual diet by selecting food stuffs from the table in Step 5, 6, 7, 8 and 9 on the following pages.

The food stuffs must not only provide the desired dietary components but must also be mixed to fit the caloric specifications. Be aware of the relative content of protein in the diet as well as the highly variable content of the protein in food. It is advisable to balance dietary selection with protein in food. The CHO, fat and total allowance will affect the protein values of the various food stuffs must be kept in mind.

STEP 5 - DETERMINE THE PROTEIN NEEDS AND SELECT FOODS TO BE USED

Protein are necessary to go with development this is especially true of energy. On Gm. of protein 4 Calories 100 Gm. of protein during its metabolism may yield about 58 Gm. CHO.

RECOMMENDED DAILY PROTEIN ALLOWANCES (N R C 1953)

	Amount per unit of body weight	
1 Adolescent	1.5-2.0 Gm./Kg.	0.7-0.9 Gm./lb.
2 Adult	1.0 Gm./Kg.	0.5 Gm./lb.
3 Pregnant woman	1.5 Gm./Kg.	0.7 Gm./lb.
4 Lactating woman	2.0 Gm./Kg.	0.9 Gm./lb.

Most of the protein requirements will be obtained from high protein foods which form the basis of the protein intake. After determining the amount of protein needed for the diet select the high protein foods by the use of the table on the following page.

HIGH PROTEIN FOODS

These proteins are interchangeable in the diet. One serving yields about 6 Gm of protein; however, the total caloric content varies.

Food	Serving	Protein		CHO	Fat	Total
		Gm	Cal	Gm	Gm	Cal
Egg	1 small	6	24	0	6	75
Milk skimmed	1 c p or glass	6	24	10	0.6	65
Milk whole	(200 cc)	8	24	10	8	130
Lean meat or fish	1 oz (fresh)	6	24	0	5	70
Fatty meat or fish	(30 Gm)	6	24	0	7	90
Fresh fowl	1 oz (30 Gm)	8	24	0	2	40
Cottage cheese	1 round d Tbsp.	6	24	1	0	30
Processed cheese	1 slice (1 oz.)	8	24	0.5	8	100
French soy beans	1/4 cup	6	24	2	3	60
Other legumes	1/2 cup	8	24	1.5	1	100
Nuts	1 oz	8	24	2	16	200

RELATIVE PROTEIN VALUES OF PROTEIN PORTION OF DIETS

Diets of more than 70 Gm or less than 40 Gm of protein can be calculated by either adding or dividing the basic portions. The table below is so arranged that proteins for low caloric (low fat) and normal or high caloric diets can be selected.

PROTEIN PORTIONS OF DIETS

For Low caloric (Low fat) Diet		For Normal or High caloric Diet	
Cal		Cal	
Yields 40 Gm or 160 Cal Protein		Yields 60 Gm or 200 Cal Protein	
1 Egg	75	1 Egg	75
2 Cups skim milk (400 cc)	130	2 Cups whole milk (400 cc)	280
3 1/2 oz meat (lean)	245	3 1/2 oz meat (med fat)	315
	40		650
Yields 50 Gm or 200 Cal Protein		Yields 60 Gm or 240 Cal Protein	
1 Egg	75	1 Egg	75
2 Cups skim milk (400 cc)	130	2 Cups whole milk (400 cc)	280
2 Tbsp cottage cheese	60	2 Tbsp cottage cheese	60
3 1/2 oz meat (lean)	245	3 1/2 oz meat (med fat)	315
	510		710
Yields 60 Gm or 240 Cal Protein		Yields 70 Gm or 280 Cal Protein	
1 Egg	75	1 Egg	75
2 Cups skim milk (400 cc)	130	2 Cups whole milk (400 cc)	280
1/2 Cup cottage cheese	240	1/2 Cup cottage cheese	240
3 1/2 oz meat (lean)	245	3 1/2 oz meat (med fat)	315
	690		890
Yields 70 Gm or 280 Cal Protein		Yields 80 Gm or 320 Cal Protein	
1 Egg	75	1 Egg	75
2 Cups skim milk (400 cc)	130	2 Cups whole milk (400 cc)	280
1/2 Cup cottage cheese	240	1/2 Cup cottage cheese	240
5 oz meat (lean)	350	5 oz meat (med fat)	450
	795		1025

*Total calories represent the caloric value derived from the carbohydrate, fat, and protein content of the foods listed.

STEP 6 - SELECT THE CARBOHYDRATE FOODS FOR THE DIET

Carbohydrates supply energy and usually constitute the largest part of the diet (about 50%). One Gm. of CHO = 4 Cal. If desired, CHO can give the proteins are separated as sources of energy. At least 10-15% of the diet must be CHO to prevent ketosis.

A. For rough approximation of the CHO content of foods, the following figures will suffice. An average serving is approximately $\frac{1}{2}$ cup cooked or 1 cup raw vegetables or fruits.

Average Serving	Amount of CHO	Total Calorie
Vegetable	4-8 Gm.	25
Fruit	12-15 Gm.	50
Shelb ead potatoes		
corn beans c eals	15-20 Gm.	75

B. For close approximation of the CHO content of food

1. 5% vegetables and fruits: 100 Gm. portion yields 3-7 Gm. CHO, 1 Gm. protein and approximately 5 Calories.

Asparagus (8 stalks)	Cucumbers (20 slices)	Spinach (1 c)
Bamboo shoots (3/4 c)	Eggplant (2 slices)	String bean (1 c)
Bananas (1 lb)	Endive (1 bunch)	Summer squash (1 lb)
Beet greens (1 c)	Lettuce (1/3 bunch)	Tomatoes (1 small)
Boccoli (1 c)	Mustard greens (1 c)	Turkey greens (1 c)
Cabbage (1 1/2 lb)	Okra (10 pods)	Cantaloupe (1/4)
Cauliflower (1 lb)	Paper greens (1 lb)	Rhubarb (1 lb)
Celery (5 stalks)	Rice (15)	Strawberries (12)
Chard (1 1/2 c)	Sauerkraut (2/3 c)	Watermelon (1/2 slice)

2. 10% vegetables and fruits: 100 Gm. portion yields 8-12 Gm. CHO, 1 Gm. protein and approximately 40 Calories.

Artichoke (1)	Onions (2)	Gooseberries (2/3 c)
Beet (2/3 lb)	Pumpkin (1/2 c)	Grapefruit (1/2 c juiced)
Carrots (1 large)	Rutabaga (3/4 lb)	Honeydew melon (1/10)
Dandelion greens (1 c)	Whiteturkey (3/4 lb)	Orange (1 juiced)
Green bean (1 lb)	Winter squash (1 lb)	Peach (1 large)
Lentils (4 stalks)	Cranberries (1 lb)	Tangerines (2)

3. 15% grains and fruits (for breakfast and without vegetables): 100 Gm. portion yields 12-17 Gm. CHO, 1 Gm. protein and approximately 50 Calories.

Apples (1 medium)	Corn (1 lb)	Peas (3/4 lb)
Apricots (2)	Grape (1 c)	Pineapple (2 slices)
Blackberries (1 lb)	Loganberries (1 lb)	Plum (3)
Blackberries (2/3 lb)	Nut (1 lb)	Raspberries (1 lb)
Cherries (1 lb)	Pear (1 lb)	

4. High CHO food: Serving yields approximately 75-100 Calories. 1/2 cup of all macaroni, gumbo (1 slice), 3-5 Gm. protein.

b. 1/2 cup of pumpkin and potatoes (also yields 1 Gm. protein). 1 slice of bread (also yields 2 Gm. protein).

d. Chickpeas, lentils (also yields 2 Gm. protein). 4 oz. 3 g. h. m. 5 p. t. is 3 Ry. k. isp.

Did fruit 1/4 p. in 3 o. 4 l. rg. p. e. or dat. 1 2 l. g. figs (also yields 1 Gm. protein).

f. Sugar 3 b. o. 2 heaping spoonful.

52 Fat and Vitamins

C B e g (Cal / 30 cc) (For milk see page 50) Black coffee
1 tea 0 ginger ale 12 beer 12 other carbonated be e ag s
15 dry win 25 sweet wine 40 liq ors 75 (Caloric values
for be r wine and liquo s are derived mainly from alcohol)

STEP 7 - DETERMINE THE FAT REQUIREMENTS AND THE DIETARY SOURCES

The requirements for fat are not known but fat forms an important source of food. 1 Gm fat = 9 Cal. Fats usually make up the remainder of the caloric intake after protein and CHO portions have been selected. Most protein-containing foods contain fat also; this must be calculated in determining total fat intake (see page 50).

The role of essential (unsaturated) fatty acids has recently been reemphasized. It has been shown that using 50-150 Gm per day of oil containing high concentrations of essential fatty acids plasma cholesterol can be lowered. The exact amounts of essential fatty acids needed and their precise role in human nutrition or human diseases are still undetermined. However, the average diet should probably contain essential fatty acids in adequate amounts. Some of the common edible oils and their average essential fatty acid (linoleic acid) content are as follows: safflower oil 70% corn oil 42% cottonseed oil 45% soy oil 54% olive oil 7%.

Caloric Values of Servings of Fats

Each of the following quantities about 40 Calories: 1 tsp butter, 1 tsp lard, 1 tsp margarine, 1 tsp mayonaisse, 1 tsp animal fat, 1 Tbsp light cream, 1 tsp oil, 1 strip bacon. (One square of butter or margarine equals 80-100 Calories.)

STEP 8 - DETERMINE THE NEED FOR VITAMIN SUPPLEMENTS

The normal daily requirements are adequately supplied by the basic diet shown on page 47. It is only in cases of restricted diet or harmful metabolic states (e.g., diabetes, fever, thyrotoxicosis, digestive absorption defect) that vitamin supplements may be necessary. For the appropriate dosages see pages 60 to 64.

DAILY ALLOWANCES OF VITAMINS (N R C 1953)

Vitamin and Daily Requirement	Natural Sources
A 5,000-8,000 I.U.	Vitamin A: Milk, butter, and liver oil; Carotene: prunes, carrots, sweet potatoes, apricots, spinach, green leafy vegetables.
B ₁ Thiamin 1.2-1.6 mg	Yeast, malt, whole grain, bran, live yeast, egg yolk.
B ₂ Riboflavin 1.4-2.5 mg	Milk, yeast, eggs, liver, meat.
P.P. Nicotin 10-16 mg	Liver, yeast, meat, bran, whole wheat.
C Ascorbic Acid 70-150 mg	Citrus fruits, green peppers, parsley, tomatoes, cabbage, radishes.
D 400 U	Battle liver, yellow fish liver oil.

STEP 9 - DETERMINE THE NEED FOR MINERAL SUPPLEMENTS

Daily requirements of the minerals are supplied in a well balanced diet (see page 47). Additional amounts are required when an abnormal loss or increased demand arises. The given mineral is usually the one prescribed as a drug. The deficiencies most likely to occur are those of calcium and iron. Iodine deficiency in endemic areas can be prevented if iodine salt is used.

DAILY ALLOWANCES OF MINERALS (N R C 1953)

Mineral	Allowances	Nutritional Sources
Mineral requirement to be deficient		
Calcium	0.8 Gm for adults 1.5 Gm for pregnant and lactating women	Milk and milk products (1 Gm calcium/qt)
Iron	12-15 mg for children menstruating women Less than 10 mg for infants	Liver, egg yolk, kidney, beef, whole wheat, green vegetables
Mineral requirement to be deficient		
Copper	1-2 mg	Liver, egg yolk, bran, oatmeal
Iodine	0.12-0.3 mg	Iodized salt, Veg table, seaweed, iodine, kelp, goit
Sodium	2-5 Gm	Table salt, milk, meat, eggs
Phosphorus	1-1.5 Gm (2-5 Gm during pregnancy)	Milk, liver, egg yolk, cereals, nut, beans
Potassium	1-4 Gm	All fruits, vegetables and fruits

STEP 10 - ARRANGE THE NUMBER, FREQUENCY AND TIME OF FEEDINGS

In the management of vertebrate animals, it may be common to vary the giving of a certain number of feedings in the life of a species at different intervals as one of the greater portions of the day. The conditions in which the animal is kept are important factors.

1. Malnutrition of the vertebrate is usually a result of frequent giving of food to the animal.
2. Frequent feeding of the vertebrate is usually a result of frequent feeding of the animal.
3. Frequent feeding of the vertebrate is usually a result of frequent feeding of the animal.
4. Frequent feeding of the vertebrate is usually a result of frequent feeding of the animal.

STEP 11 - GIVE DETAILED INSTRUCTIONS TO THE PATIENT

When the diet has been completely planned, carefully explain it to the patient in a simple and direct manner. The patient should be told the position of the feedings, the number of meals and the time of feeding. The following explanation of the diet will aid in the understanding of the instructions.

PRINCIPAL TYPES OF DIETS

The following diets are planned around the Basic Foods which form the nucleus of a well balanced diet. See table on page 47.

Sippy Diets

Progressive non irritating buffering diets taken on regular schedule

Composition

- Stage I 3 oz (90 cc) half milk and half cream (18%) every hour from 7:00 a.m. to 7:00 p.m.
- Stage II Stage I plus 3 feedings of refined cereal (3 pe serving) and 1 soft cooked egg tid
- Stage III Stage II plus creamed soups and pureed vegetable
- Stage IV 3 (90 cc) milk and cream every hour plus regular meal of small feedings of lean meat, potato, pureed vegetable, refined cereals and breads, custard puddings, cream and butter.

Restrictions Meat extracts, bran, raw vegetable and fruits, tea, free condiments, pepper, alcohol and alcoholic beverages.

Mulengah's Diet

Employed in bleeding patients. As now generally employed means frequent feedings of pureed foods. Originally described as follows:

- 6 a.m. Tea, white bread and butter
- 9 a.m. Oatmeal with milk, white bread and butter
- 1 p.m. Dinner. A muhasa, red of meat, broiled chops, omelet, fish, vegetable, cream at or fl, h gratin, mashed potatoes, vegetable puree, or soups, creamed vegetable, stewed apples, apple sauce, gruel, and rice and tapioca puddings.
- 3 p.m. Cocoa
- 6 p.m. White bread and butter, sliced meats, cheese and tea.

Bland Diet

An ideal diet modified to be suitable for non irritating and bland intake. May also be used as a low residue diet.

Composition Lean meats, fish, poultry, egg, milk, potato, pureed vegetable and fruit, refined cereals and bread, custards, puddings, gelatin desserts, cream, butter, margarine, salt and sugar in moderation.

Restrictions Fried food, raw vegetables, fruits and fruit juices, spice, condiment, bran, whole grain cereals and bread, carbonated beverages, alcohol and coffee.

Low Fat No Gas Formula Diet

Composition Lean meat, fish, poultry, skimmed milk, butter, milk, cottage cheese, cereal products, bread, vegetables and fruits except those listed below, gelatin desserts, sherbet, pudding without cream, sugars and jellies.

Restrictions Pork, ham, bacon, fatty cuts of any meat, cream, cabbage, family onions, turnip, cucumber, radishes, green peppers, dried beans and peas, molasses, raw apples, butter, margarine, mayonnaise, oil, nuts, chocolate, fried foods, pastries and highly seasoned foods.

High prot i High CHO Low fat Diet

Composition All w fat d t w th r ess placed o la ge se ings
of l an m t gg skimmed milk o buttermilk cottage
che se c al b ada f uit jui sug r and jelly To c l
cul te a d f nite amount of p ot in fo thi diet see tabl s on
p ge 50

Restrictions S m a f r low f t non g s forming di t

High resid e Diet

A normal diet with a maximum of bulk

Composition All of th ba i foods with ext rvings of whole
gr in ls and breads r w egetabl a d fruits a d an
adequate mount of fluids

Restriction Non

Diet Restricted in Sodium Content

S d um r et t d diets u ally mploy 1.5 Gm. of sod um
(3.75 Gm. od um chl de) or und r Fo best th peptic r
sult diets should conta n less th n 500 mg. of sod um (1.5 Gm.
sod m hlo d)

Th f llowing tw low sod um diets both cont in 2 000 C lories
Th y r th sam n comp sition c pt for the beverag

250 mg. sod um di t u e Lonala ® a bev g

500 mg. sod um diet us wh le milk a bever g

Breakfast

Fruit	1 ¹ / ₂ c p
S lt f e co k d r puff d c al	1 ¹ / ₂ c p
Salt f e b d	1 slice
S lt f e b tt m g in	2 t p (1 pat)
Egg	1
Lon l c® or whol milk	1 ¹ / ₂ p

Noon and Evening Meals

S lt f e l sh m t	3 ¹ / ₂ o
S lt f e pot to o i	1 ¹ / ₂ up
Salt f e o ked o r w veget bles	as d ir d
S lt f bre d	1 slice
S lt f e butte or ma g ine	2 tsp (1 p t)
F ut	1 ¹ / ₂ up
L al c® or whole milk	8 oz

Additional (S Restriction on Following Page)

- 1 Lo al ® is p p r d by m ing 1¹/₂ up d y p wd w th 2
ups of w t th sm y b fl o d with ho lat
- 2 T m k lt fr m g in w h and knead m g in in
flv hange of old w t
- 3 U t h o f oz n v g t bl sp l lt fr n n d
g t bl M y u rt h ke be t ote pin h and
oth g twl weekly
- 4 U nly f h o k d f ut
- 5 U o ly granu l d g l tin in salad and d t
- 6 M y u e p p h b e and othe spi
- 7 M y u one of th od m f salt ub titut e

B Restrictions

- 1 Ham bacon bacon fat salt pork corned beef or pork luncheon meats canned meats fish or poultry
- 2 Prepared cereals with salt quick cooking cereals breads leavened with baking powder or baking soda
- 3 Prepared foods or prepared desserts
- 4 Canned vegetables dried fruits commercial salad dressings catsup
- 5 Salted nuts salted popcorn potato chips
- 6 Garlic salt onion salt celery salt salt baking powder
- 7 Celery olives pickles fishes chard
- 8 To avoid distention Cabbage family onions turnips peppers dried bean cucumbers sweet potatoes raw apples melons

C Approximate Sodium Content of Common Foods (in mg per serving) This list gives the natural content without the addition of salt baking powder or baking soda

- 1 Fresh meat fish and poultry $3\frac{1}{2}$ oz (100 Gm) serving

Lamb	78	Oyster	73	Chicken leg	110
Pork	59	Cod fish	60	Turkey leg	92
Beef	51	Hamburger	56	Chicken breast	78
Veal	48	Salmon	48	Turkey breast	40
- 2 Egg (1) 40
- 3 Milk 7 oz glass (200 cc) Cultured buttermilk 270 fresh whole milk 110 reconstituted whole milk (Longlac®) 3
- 4 Cheese 1 oz (30 Gm) Processed 450 cheddar 210 cottage 100 cream 75
- 5 Legumes $\frac{1}{2}$ cup (4 oz or 120 Gm) fresh or $\frac{1}{8}$ cup (1 oz or 30 Gm) dry Beans and corn 0 split peas dry 42
- 6 Cereals 1 oz (30 Gm) dry $\frac{1}{4}$ cup whole grain cereals or pastes (macaroni etc) 0 5 4 1 cup dry cold cereals 200 350 puffed cereals 1
- 7 Bread (1 slice) and crackers

Commercial bread	180 250	Sanitary crackers	230
Yeast bread without salt	0 5	Malt loaf plain	0 3
- 8 Vegetables $3\frac{1}{2}$ oz (100 Gm) serving of fresh or frozen (not canned) (For sizes of serving see pag 51)

Artichoke	43	Cabbage	5	Endive	18	Potatoes	
Asparagus	2	Carrots	31	Kale	110	Skim milk	0 6
Beans	1 2	Cauliflower	34	Lettuce	12	Pumpkin	0 4
frozen	2	Chard	200	Okra pods	1	Spinach	82
Beets	110	Celery	110	Onion	1	Squash	0 5
Broccoli	16	Corn	trace	Parsnips	7	Tomato	3
Brussels		frozen	9	Peas	0 9	Turnip	37
sprouts	16	Eggplant	1	French	100		
- 9 Fruits $3\frac{1}{2}$ oz (100 Gm) serving (if fresh see pag 51)

Fresh canned and frozen fruits contain less than 10 mg sodium per serving					
---------------------------------------------------------------------------	--	--	--	--	--
- 10 Fats 10 Gm (2 teaspoons)

Margarine	110	Sweet butter	0 5	Shortening	0 1
Regular butter	98	Oil	0 2	Lard	0 3
- 11 Sweets 10 Gm (2 teaspoons)

Sugar	minute amounts	honey	2 0	jelly	0 2
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1 Miscellaneous
 Breakfast 19 Coca Cola® 1 bottle 18
 Cereal 19 Nuts 1 oz (30 Gm) 0.5
 Coffee, tea, natural herbs and condiments only
 negligible amounts of sodium

Diabetic Diet

A calculated diet with calculated amounts of protein, fat, and carbohydrate

1800 CaloBreakfast (7:00-9:00 a.m.)

1/2 cup 10% fruit
 2 eggs any style
 1 tsp butter or margarine
 1 glass skimmed milk

Morning Feeding (10:00 a.m.)

1 glass skimmed milk
 1 inch cube processed
 cheddar cheese

Noon Meal (12:00-1:00 p.m.)

3 oz any lean meat
 chicken or fish
 1/2 cup 5% vegetable
 1/3 cup 5% vegetable
 salad 5% vegetable
 2 tsp butter or margarine
 1/2 cup 10% fruit
 1 glass milk

Afternoon Feeding (3:00 p.m.)

1 glass skimmed milk
 1 ounce dried
 peanut butter

Evening Meal (6:00-7:00 p.m.)

3 oz any lean meat
 chicken or fish
 1/2 cup 5% vegetable
 1/3 cup 5% vegetable
 salad 5% vegetable
 2 tsp butter or margarine
 1/2 cup 10% fruit
 1 glass milk

Bedtime Feeding (9:00-10:00 p.m.)

1 glass skimmed milk
 1/2 cup (any) cottage
 cheese

2500 CalorieBreakfast (7:00-9:00 a.m.)

1/2 cup 10% fruit
 2 eggs any style
 2 strips bacon (1 p)
 Coffee or tea as desired

Morning Feeding (10:00 a.m.)

1 cup whole milk
 2 inch cubes processed
 cheese or 1/3 cup peanuts

Noon Meal (12:00-1:00 p.m.)

1/2 cup processed cheese
 1/2 cup 5% vegetable
 1/2 cup 10% vegetable
 1/2 cup 10% fruit
 2 tsp butter or margarine
 1 cup whole milk
 Coffee or tea as desired

Afternoon Feeding (3:00 p.m.)

1 cup whole milk
 2 inch cubes processed
 cheese or 1/3 cup peanuts

Evening Meal (6:00-7:00 p.m.)

4 oz lean meat chicken
 fish
 1/2 cup 5% vegetable
 1/2 cup 10% vegetable
 1/2 cup 10% fruit
 2 tsp butter or margarine
 1 cup skimmed milk
 Coffee or tea as desired

Bedtime Feeding (9:00-10:00 p.m.)

1 cup whole milk
 2 inch cubes processed
 cheese or 1/3 cup peanuts

58 Diets

High calorie High protein High vitamin Diet

A normal diet containing extra foods high in protein and all of the vitamins

Composition All of the basic foods with increased amounts of meat fish poultry liver eggs milk cheese whole grain cereals carrots green vegetables citrus fruit butter or margarine (see table on page 58 for high protein foods and table on page 52 for high vitamin foods)

Restrictions None

Low calorie Diets

Basic diets containing adequate protein which are low in calories than the patient's daily requirement (see page 47). The amount of food listed in each diet is the total daily intake

1200 Calorie Diet

5 oz meat fish poultry or cheese

1 egg

1 pt skimmed milk or buttermilk

1 slice bread

1 serving (1½ cup) potato equivalent

2 servings 10% vegetables

3-4 servings 5% vegetables

2 servings 10% fresh fruit

1 serving 15% fresh fruit

2 tsp butter or margarine

1000 Calorie Diet

Omit the following from the 1200 Calorie diet

1 serving potato 1 serving 5% vegetables

1 serving 10% vegetable 1 tsp butter

800 Calorie Diet

3 oz lean meat fish or poultry

2 oz cottage cheese

1 slice bread

1 serving 10% vegetables

3 servings 5% vegetables

3 servings 10% fruit

1 pt skimmed milk or buttermilk

1500 Calorie Diet

Add the following to the 1200 Calorie diet

2 slices bread

3 tsp butter or margarine

1 serving 15% fruit

Restrictions All foods candy and beverages except those listed

Low protein Diet

A normal diet with the protein foods limited to the minimum biologically adequate amount

Special Elimination Diets

A normal diet containing no foods suspected of causing allergic reactions. Such reactions reproduced most frequently by wheat, eggs and milk less frequently by citrus fruits, chocolate and fish. Other foods may infrequently cause reactions.

More specialized diets have been prepared by allergists and are used both diagnostically and therapeutically. Consult books on allergy for these diets.

Low purin Diet

Diet low in protein

Food Forbidden Liver kidney sweetb ads sardines anchovies
 beans whole grain products gravy soup meat extracts
 asparagus bean cauliflower peas lentils and mushrooms

Food Restricted All other meat fish and fowl

Composition All other foods are allowed. Most protein to be
 derived from eggs and dairy products

TUBE FEEDINGS

Tube feeding is employed when patient is unable to swallow
 to take food by mouth. A convenient means of administering the
 feeding is with a small polyethylene tube passed into the stomach.
 Many food mixtures may be given throughly equipped with the
 the food be fluid or in a suspension of very small particles.

Protein hydrolytes are often irritating. Formula containing
 egg yolk to occlude the lumina of small intestines. Excellent formulas
 can be prepared by using milk (occasionally hot in tube) calcium
 caseinate. Lactogen[®] is tried in infants. Use of glucose
 formula such as Isodol may be added if emulsified with Tween 80[®]
 as emulsifier. Vitamins and minerals are added as indicated.
 Example of tube feeding formula is as follows

- 1 Low lipid high protein diet. Supply 3 000 Calories
 per 3 000 c (1 Cal /) contains 133 Gm protein
 Sterilized condensed milk 400 Gm (4 cans)
 Tomato juice 1900 c
 Prun juice 90 c
 All purpose Soyabean[®] 200 Gm
 Lactogen 315 Gm (1 1/3 cup)
 Water 3000 c
- 2 Experimental high protein formula. 3 000 Calories per
 3 000 c (1 Cal /) contains 120 Gm protein
 Homogenized milk 2200
 1/2 milk and 1/2 cream 600 c
 Eggs 6
 Dextrose[®] lactose 7 Tbsp
- 3 Low sodium high protein formula. 3 000 Calories per
 3 000 c contains 150 Gm protein 78 mg sodium
 Lactogen 600 Gm
 Water 3000 c

THE VITAMINS

The vitamins are organic substances which
 must be supplied to the organism for growth and health. They
 are essential for the proper functioning of the body. They
 are important for the health of the body. They are essential for
 the health of the body. They are essential for the health of the
 body. They are essential for the health of the body. They are
 essential for the health of the body. They are essential for the
 health of the body. They are essential for the health of the body.

58 Diets

High calorie High protein High Vitamin Diet

A normal diet containing extra foods high in protein and all of the vitamins

Composition All of the basic foods with increased amounts of meat fish poultry live eggs milk cheese whole grain cereals carrots green vegetables citrus fruits butter or margarine (see table on page 58 for high protein foods and table on page 52 for high vitamin foods)

Restrictions None

Low calorie Diet

Bulky diets containing adequate protein which are lower in calories than the patient's daily requirement (see page 47) The amount of food listed in each diet is the total daily intake

1200 Calorie Diet

5 oz meat fish poultry or cheese
1 egg
1 pt skimmed milk or buttermilk
1 slice bread
1 serving ($\frac{1}{2}$ cup) potato or equiv lent
2 servings 10^m vegetable
3-4 servings 5^m vegetable
2 servings 10^m fresh fruit
1 serving 15% fresh fruit
2 tsp butter or margarine

1000 Calorie Diet

Omit the following from the 1200 Calorie diet

1 serving potato 1 serving 5^m vegetable
1 serving 10^m vegetable 1 tsp butter

800 Calorie Diet

3 oz lean meat fish or poultry
2 oz cottage cheese
1 slice bread
1 serving 10^m vegetable
3 servings 5^m vegetable
3 servings 10^m fruit
1 pt skimmed milk or buttermilk

1500 Calorie Diet

Add the following to the 1200 Calorie diet
2 slices bread
3 tsp butter or margarine
1 serving 15% fruit

Restriction All foods candy and beverages except the selected

Low protein Diet

A normal diet with the protein foods limited to the minimum but adequate amount

Sensible Elimination Diet

A normal diet containing all foods excepted of single allergenic reaction. Such reactions are produced most frequently by wheat egg and milk less frequently by citrus fruits chocolate and fish. Other foods may frequently cause reactions.

No specialized diets have been prepared by all allergists and are used both diagnostically and therapeutically. Combination of allergy for these diets

Vitamin D (Code No 010 754)

Avitaminosis D is usually due to deficiency of sunlight or absorption defect

A Child rickets Lack of vitamin D leads to rickets in child
 Child is known as rickets (see page 380)

B Tetosis Deficiency Serum calcium and phosphorus may be normal or decreased and alkaline phosphatase is generally increased
 Increased urinary calcium excretion; decreased

C Treatment See Osteomalacia page 380

VITAMIN E

Vitamin E plays a role in normal physiology of some animals but there is no good evidence of its activity in man. It is relatively toxic. It has been used without apparent benefit in some cases of habitual abortion, varicose veins, muscular dystrophies and heart disease. Dose of 30-100 mg of tocopherol daily.

VITAMIN K

The vitamins K are chemical compounds necessary for proper blood synthesis by the liver and are important in the blood coagulation mechanism. They are widely distributed in all vegetables, fruits and animal products. They are also synthesized by microorganisms in the intestine. All water soluble

Vitamin K (Code No 010 755)

A Avitaminosis K Its formation depends on synthesis of prothrombin, deficiency leads to prothrombin deficiency (coumarin sensitivity) drug dependence of the blood synthesis (coumarin sensitivity)

A Meningitis Bleeding from mucous membranes of the mouth

B Tetosis Deficiency Prolonged prothrombin time

C Treatment

1 Pre-treatment fluids electrolytes pp 279-287

2 Pre-treatment fluids electrolytes pp 216

3 Pregnancy fluids electrolytes 2.5 mg M dose
Sodium B-sulfate injection USP (Mephathel injection)
BP IV IM 172 hours before delivery pre
bleeding newborn infants

WATER SOLUBLE VITAMINS

VITAMIN B COMPLEX

The members of the vitamin B complex are very intimately related to each other in function. A deficiency of one of the members of the complex leads to deficiency of all the members. The members of the B complex would lead to impairment of the metabolism of the other. However, although certain clinical features may point to the deficiency of one of the members of the complex, this does not signify that the deficiency can be truly corrected by administering the single factor. Therefore, the proper

In illness there may be considerable variation in the body requirements depending upon age, activity, diet, metabolic rate, and other factors affecting the absorption, utilization, and excretion of vitamins. Vitamin deficiencies are almost always multiple, particularly of fat-soluble or B-complex vitamins as a group. Early signs of vitamin deficiency are usually nonspecific, vague, and mild and are easily misinterpreted or missed entirely. The crude sources of the vitamins are often more efficacious in therapy than the pure or synthetic. Only during the most severe phases of the deficiencies is it usually necessary to resort to the use of pure vitamins. The use of a pure vitamin in the face of a true multiple vitamin deficiency may aggravate rather than help the condition. Treatment of vitamin deficiencies requires an adequate, balanced, high-protein and high-vitamin diet in addition to necessary vitamin supplements. In general, it is wise to use vitamins therapeutically in 5-10 times the amount required for daily maintenance.

FAT-SOLUBLE VITAMINS

VITAMIN A

Vitamin A is an alcohol of high molecular weight which is converted from β -carotene in foods by the liver. It is necessary for normal function and structure of all epithelial cells and for synthesis of visual purple in retinal rods (necessary for vision in dim light). It is present in leafy green and yellow fruits and vegetables, whole milk, butter, eggs. Recommended daily allowances for adults are 5000 I.U. (60 U.S.P. units) during pregnancy and lactation, 6000 to 8000 I.U. Toxicity: 500,000 to 1,000,000 I.U. daily may cause alopecia, itching, and bone pain from periosteal proliferation.

Vitamin A (Code No. 010-761)

- A Mild or Early Manifestations: Dryness of the skin, night blindness, and follicular hyperkeratosis.
- B Severe or Late Manifestations: Xerophthalmia, atrophy and keratinization of the skin, and keratomalacia.
- C Tests for Deficiency: Dark adaptation is impaired. Low blood levels of vitamin A may be helpful or a therapeutic trial with 25,000 to 75,000 I.U. daily for 4 weeks.
- D Treatment: Of vitamin A, U.S.P. Vitamin A, B.P. 15,000 to 25,000 I.U. or 600 to 1,000 units twice daily. If absorption defective, it may be necessary to administer bile salts with the vitamin A or to give the same dosage in oil (1 M (50,000 units/cc in sesame oil). Skin lesions may require moist emollients.

VITAMIN D

The vitamin D are sterols formed in the animal body by ultraviolet irradiation of plant sterol precursors. They increase calcium absorption from the intestine and urinary phosphorus excretion. They are present in fish livers, their precursors are widely distributed in plants. Allowances for adults are not known. For children and during pregnancy and lactation, 400 I.U. (or U.S.P. units) are recommended. Toxicity: 150,000 I.U. or more daily causes elevated serum calcium with metastatic calcification.

Avitaminosis D (Cod No 010 764)

Avitaminosis D is usually due to inadequate dietary intake lack of night or absorption defect

A Clinical Finding Lack of vitamin D leads to osteomalacia in children this is known as rickets (see page 380)

B Test for Deficiency Serum calcium and phosphorus may be normal or decreased and alkaline phosphatase is generally increased urinary calcium excretion is decreased

C Treatment See Osteomalacia page 380

VITAMIN E

Vitamin E plays a role in the normal physiology of some animals but there is no good evidence of its activity in man. It is relatively toxic. It has been used without apparent benefit in some cases of heart labilation via a neuromuscular syndromes and heart disease and doses of 50-100 mg of tocopherol daily

VITAMIN K

The vitamin K chemical compounds are say for example menin synthase by which the carboxyl group is important in blood coagulation mechanism. They are widely distributed in green leafy plants egg yolk and soybeans. They are also synthesized by microorganisms in the intestines. Allowance is 5-10 mg daily

Avitaminosis K (Cod No 010 766)

Avitaminosis K is from liver disease interfering with synthesis of prothrombin and adequate bile supply with poor absorption of digested prothrombin through the intestine (carboxyl salts)

A Mechanism Bleeding from microvascular trauma

B Test for Deficiency Prolonged prothrombin time

C Treatment

1 Factorial treatment see pp 279-287

2 Factorial treatment see pp 216

3 Pregnant women give daily 10 mg of vitamin K (Mephyton Inj 10 mg/ml B.P.) or 1 mg 12-22 hours before delivery to prevent bleeding in newborn infants

WATER-SOLUBLE VITAMINS

VITAMIN B COMPLEX

The members of the vitamin B complex are very intimately associated in their chemical functions. As a result of this it is difficult to separate them. It is doubtful that a deficiency of a single B vitamin is a distinct entity and the clinical picture of a deficiency of a single member of the B complex would be indistinguishable from that of a deficiency of the whole complex. This does not signify that the deficiency is not biologically distinct but admits that the single deficiency is not the same as the deficiency of the whole complex.

must always be supplied in the presence of adequate (dietary or parenteral) sources of all of the other members of the B complex. Water soluble vitamins should be administered in divided doses throughout the day to prevent excessive loss in the urine.

VITAMIN B₁ (Thiamine Hydrochloride)

Vitamin B₁ is the coenzyme for decarboxylation of α keto acids (e.g. pyruvic & ketoglutaric). It is important therefore for normal carbohydrate oxidation. Dietary sources are liver, lean pork, kidney and whole grain cereals. Steaming or exposure to moist heat reduces the thiamine content of foods. Daily dietary allowances are about 0.5 mg per 1000 Calories (avg. 1.2-1.6 mg per day).

Avitaminosis B₁ (Beriberi) (Code No. 010 7621)

Avitaminosis B₁ results from an inadequate intake due usually to idiosyncrasies of diet or excessive cooking or processing of foods. The increased need for vitamin B₁ during fever, high CHO intake or thyrotoxicosis may lead to a deficiency.

A Mild or Early Manifestations Vague multiple complaints suggest nerve disease and include anorexia, formication and muscle cramps, tenderness of calves, paraesthesiae and hyperactivity followed later by hypoactivity of knee and ankle jerks.

B Severe or Late Manifestations (Beriberi) Severe anorexia, polyneuritis, serous effusions, subcutaneous edema, paralysis (particularly in extensors) and cardiac insufficiency manifested by tachycardia, dyspnea, edema and normal or decreased circulation time, elevated venous pressure and non-specific ECG changes.

C Treatment

1. Thiamine Hydrochloride U.S.P. (Aneurin Hydrochloride B.P.) 20-50 mg orally I.V. or I.M. divided in divided doses for 2 weeks then 10 mg daily orally.
2. Dried Yeast Tablets U.S.P. (brewer's yeast) 30 Gm t.i.d.
3. Well balanced (2500-4500 Calorie) diet when tolerated.

VITAMIN B₂ (Riboflavin)

Riboflavin serves as coenzyme for hydrogen transfer. It is present in milk and milk products, leafy green vegetables and liver. Daily dietary allowances for adults are 1.4-1.8 mg in pregnancy and lactation 2.2-5 mg.

Avitaminosis B₂ (Ariboflavinosis) (Code No. 010 7622)

The biological factors of ariboflavinosis are similar to those of thiamine deficiency but inadequate intake of milk is important. The manifestations of deficiency usually occur along with thiamin and niacin deficiency but may occur earlier.

A Mild or Early Manifestation Oral plaques, superficial fissuring at angles of mouth, conjunctivitis and photophobia, lack of vigor, malaise, weakness and weight loss.

B Severe or Late Manifestations Cheilosis (fissuring at the angles of the mouth), fissuring of the nares, magenta tongue.

with moderate dysphagia corneal vascularization and
irritation and eczematous dermatitis

C Treatment

- 1 Riboflavin U S P B P 40-80 mg I V I M orally daily until all symptoms have cleared
- 2 Dried Yeast Tablet U S P (baker's yeast) 30 Gm tid
- 3 Well balanced (2500-4500 Calories) diet when tolerated

NICOTINIC ACID (Niacin) AND NICOTINAMIDE (Niacinamide)

Niacin and nicotinamide function in important enzymic systems concerned with reversible oxidation and reduction. They are present in many of the most whole grain cereals and peanuts. Daily allowance for adults is 10-15 mg for adolescents 12-19 mg. Niacin may be used therapeutically as a vasodilating agent for headache, myalgias, rheumatological disorders and edema of the labyrinth (100 mg orally daily in divided doses). Niacinamide does not possess this vasodilating effect.

Pellagra (Code No. 010-7823)

The etiological factors of deficiency are similar to those of thiamine deficiency. Niacin deficiency is the principal but not the only dietary defect in pellagra.

- A Mild or Early Manifestations Multiple vague complaints and swollen, roughened skin. Redness and hypertrophy of the papillae of the tongue.
- B Severe Late Manifestations Marked roughening of skin when exposed to light and fetid diarrhoea abdominal distention, curled tongue with atrophy of papillae, mental depression, clouding of mentality, rigidity and peculiar screaming.
- C Treatment
 - 1 Specific measures
 - a Nicotinamid U S P B P (nicotinamide) 50-500 mg I M I V or orally daily until symptoms subside
 - Nicotinamide U S P B P (injection) is less often used because of its vasodilating effect dosage is similar
 - b Supplementary vitamins Give therapeutic doses of thiamine, riboflavin and pyridoxine
 - c Dried Yeast Tablet U S P 30 Gm tid
 - 2 General measures
 - a Well balanced (2500-4500 Calories) high protein diet
 - b Symptomatically and prophylactically indicated Dementia may require constant supervision

VITAMIN C (Ascorbic Acid)

Vitamin C is concerned with formation and maintenance of intercellular supporting structures (dentine cartilage collagen bone matrix). Its biochemical action is not known. Dietary source in food citrus fruit tomatoes, peppers, bell pepper, citrus, green, etc. Ascorbic acid content of foods is markedly decreased by cooking, mashing, contact with alkali and contact with oxygen. Deficiency leads to yellowing of skin, scurvy, 70-75 mg daily during pregnancy and lactation 100-150 mg. Ascorbic acid has also been used in the treatment of certain poisons in doses of 0.5 Gm or

must always be supplied in the presence of adequate (dietary or parenteral) sources of all of the other members of the B complex. Water-soluble vitamins should be administered in divided doses throughout the day to prevent excessive loss in the urine.

VITAMIN B₁ (Thiamine Hydrochloride)

Vitamin B₁ is the coenzyme for decarboxylation of α -keto acids (e.g., pyruvic & ketoglutaric). It is important therefore for normal carbohydrate oxidation. Dietary sources are liver, lean pork, kidney, and whole grain cereals. Steaming or exposure to moist heat reduces the thiamine content of foods. Daily dietary allowances are about 0.5 mg per 1000 Calories (avg. 1.2-1.6 mg per day).

Avitaminosis B₁ (Beriberi) (Code No. 010 7621)

Avitaminosis B₁ results from an inadequate intake due usually to idiosyncrasies of diet or excessive cooking or processing of foods. The increased need for vitamin B₁ during fever, high CRO intake, or thyrotoxicosis may lead to a deficiency.

A. Mild or Early Manifestations Vague multiple complaints suggest neurasthenia and include anorexia, formication and muscle cramp, tenderness of shins, paresthesias, and hyperactivity followed later by hypoactivity of knee and ankle jerks.

B. Severe or Late Manifestations (Beriberi) Severe anorexia, polyneuritis, serous effusions, subcutaneous edema, ptyalism (particularly in extremities), and cardiac insufficiency manifested by tachycardia, dyspnea, edema, and normal or decreased circulation time, elevated venous pressure, and non-specific ECG changes.

C. Treatment

1. Thiamine Hydrochloride U.S.P. (Aneurine Hydrochloride B.P.) 20-50 mg orally I.V. 0.1 M daily in divided doses for 2 weeks, then 10 mg daily orally.
2. Dried Yeast Tablets U.S.P. (brewer's yeast) 30 Gm t.i.d.
3. Well-balanced (2500-4500 Calorie) diet when tolerated.

VITAMIN B₂ (Riboflavin)

Riboflavin serves as coenzyme for hydrogen transfer. It is present in milk and milk products, leafy green vegetables, and liver. Daily dietary allowances for adults are 1.4-1.6 mg in pregnancy and lactation, 2-2.5 mg.

Avitaminosis B₂ (Ariboflavinosis) (Code No. 010 7622)

The etiologic factors of ariboflavinosis are similar to those of thiamine deficiency, but inadequate intake of milk is important. The manifestations of deficiency usually occur along with thiamine and niacin deficiency but may occur earlier.

A. Mild or Early Manifestations Oral pallor, superficial fissuring at angle of mouth, conjunctivitis, and photophobia, lack of vigor, malaise, weakness, and weight loss.

B. Severe or Late Manifestations Cheilosis (fissuring at the angles of the mouth), fissuring of the nares, magent tongue.

Chapter 5

DISEASES OF THE SKIN

GENERAL PRINCIPLES

History and Physical Examination

A Take a chance to yourself to be patient with a kind of
B Do not get into the of of the factors in production or
aggregation of risk disease (e.g. inter-lidase senile
factor due to aberrant etc.)
D Embrace the body's face in good light

Planning the Trialmeit

The es b wld r gv scty of d mat log c g nts l g n
r l t s b t t o b th roughly f m l i with the act o s f a f w
d 23 and t e t m e t h d th t o attempt to use a g e at many

A Consider the general character of the individual skin

- 1 D y k s s lly requi e l br ating or oftening ag nts
2 M t r lly skins us lly r qu e gr s l s drying gents

B B g i n t e m n t w t h m l d i m p l r m d i e s i g e r l

- 1 A ut nflam dlesions Soothing o i ritating g nt
Car th k dlesns Stim l tng or k atolytic
ge ts

C Apply small pressure moment of inertia to single small contact sensitivity

D D n t h g m d to oft All wagent t m to ct
How di ot m dy mm d t ly if ntows d lo l
r t nde lps

Elaborate further on how to apply standards

F. When I doubt a top operation, I first try not to UNDERTREAT
the individual.

CORTICOTROPIN (ACTH) AND THE CORTISONES

I t of th t r in of limited m
to ot t pl (ACTH) r th on sm y e t d amatic
o e e a l f i g e f f t t a b l y a n g o n u o t c d m a t p
d m t t x f l a t v d e m t t s p m p h g u l p s e y h m a t o
s d d m t o m y s t Th f f t of h i d u t a f t t h a g t
h b w t h d w R l p e a r s t u c o m m d l a g a n d a f t e r
t a t m t M a t t e t h r p y m y s p p e c n t o l c e t a n o f
th d r m t e s b t s l d m I t i s m p o t a n t t o d e
s t a n d t h p h y s i o l o g a n d p h a r m a c o l o g f f t o f t h e s e d r i

h optimum b n f t f om th py (see p g 383)

Top al s of som f th tj onc y ry

tu t t d m titi and em f

64 Vitamins

more Proof of its value is lacking It is used in dosages up to 200 mg daily orally for healing wounds or ulcers or during recovery from protracted diseases (e.g. tuberculosis)

Avitaminosis C (Scurvy) (Code No. 010 763)

Scurvy is usually due to inadequate intake but may occur with increased metabolic needs

- A Mild or Early Manifestation Edema and hemorrhage fingernails porosity of dentin hyperkeratotic hair follicles
- B Severe or Late Manifestation Severe muscle cramping swelling of the joints rarefaction of bone marked bleeding tendency extravasation of blood into fascial layers anemia loss of teeth and poor wound healing
- C Tests for Deficiency Capillary resistance is reduced and x-ray of long bones may show typical changes There is loss of weight of sediment or white blood cell ascorbic acid levels
- D Treatment
- 1 Deficiency Sodium Ascorbate I prefer to U.S.P. 0.5 to 1 Gm I.V. or 1 M daily and dedose as a long as deficiency lasts Ascorbic Acid U.S.P. B.P. may begin gradually about the same dosage
 - 2 Increased demand Ascorbic Acid U.S.P. 200-300 mg per day orally

OTHER VITAMINS

Many other vitamins have been characterized Some are important in human nutrition and some are of minor importance Among them the most important with all general clinical fulfills

Pyridoxine Hydrochloride May be important in nutrition and deficiency of it is of poor prognosis () May relieve some symptoms of weakness and peripheral neuropathy and may elevate the glossitis and help to speed by riboflavin Its role (if any) in human atherosclerosis is not known Dose 10-50 mg I.V. or I.M. daily with other factors of the B complex

Choline Found in phospholipids and methyl donor and lipotropic substance and a growth factor It is found in large quantities in yeast It has been used to treat fatty livers in the liver parenchyma in human liver disease

Folic Acid (Pteroylglutamic Acid L. C. factor) Seems to be essential for the metabolism of cellular materials Effective in certain megaloblastic anemia (see p. 222)

Vitamin B₁₂ (L. C. factor) A phosphorus and cobalt containing material is isolated from purified extracts probably the active principle lacking in pernicious anemia (see p. 222)

Vitamin P (Rutin H. P. R. D. Methyl Choline) These isolated substances may reduce capillary fragility and may also reduce the threshold of response of the peripheral phintreceptor to physical irritation in human disease a questionable Dose Hesperidin methyl chalcogen 0.5-1 Gm daily orally Rutin 20-40 mg tid qid orally

Inositol Has been shown to be a lipotropic substance under certain very special conditions in some species of animal Its role in human nutrition and its use in liver disease is still entirely unclear

Miscellaneous The roles of pantoic acid and pantoic acid aminobenzoic acid and biotin in humans are undetermined

- di der d et th r py a cordingly
- 3 E t l i r r t a n t (g r o u g h c l o t h g o c p a t n a l t i a n t s) h o u l d b a i d d
- 4 B a t h i n g p r c t i c e S o a p h o u l d b e a o i d e d i n u n d i i d u a l s w t h d r y o r i r t e d s k i n S t c h b i t h m a y b e u e d (s e p r e v i o u s p a g)
- 5 N a i s h o u l d b k p t t i m m d a n d c l a n d
- 6 A o d s t e h i n g , i f p o s s i b l b e c a s s f v i c o u c y c l w h c h a n b e t b l a s h e d
- 7 U n n e s s y m e d c a t o n h o u l d b e s t o p p d s i n c e m e d c a t n i t s i f a n o f t n p d u e p i t u
- 8 A n t p r u t i c d g T h f o l l w i n g a g n t s m y b e o f b e n f i
- Cal m G l c n a t I n j e c t i U S P 10 I V
- l o w l y o d a i l y o e v e r y o t h e r d a y p
- b A n t i h t a m i n d r u g m y b e t r i e d i n r t i n c a s s i f p r i t s o f a l l g c o r u n d e r m i n e d t o l o g y F o a l i s t o f m o n o n l y u d a n t i h i s t m i n i p r e p a r i o n s p a g e 45
- E p u r p h n I n j e c t n U S P 0 25 1 0 c (4 16 m m) o f 1 1 000 s o l u t n e v e r y 4 h o u s m y b f v a l e i n a u t c a s s s p i d o f b i n g d u e t a l l g y (r t a)
- d P h n b b i l U S P 0 015 0 03 G m (1/4 1/2 g)
- b i d o q d m y p r o d u f l e d t i o n n a g t l e d r e m o t o a l l y d i s t t e d p t i t R m m b t h t b b i r t t h m l m a y p r o d e d m a t t (e l y)
- A i t h e m t h e r p y S o m d m t o l o g i s t s d v i s t h e i n j e c t i o n o f 10 c o f t h e p t n t s w h o l e v e n o u b l o o d i t o t h h p m u s i y 48 h o u s f o r 3 n j t i o
- f C i t o t o p I n j e t o n U S P S p a g 424

DERMATITIS VENENATA (Contact Dermatitis)

(code No 110 3001)

(Dermatitis Venenata Due to Plant Irritants code No 110 378)

An cut o h d m a t t i s w h i h r u l t s f o m d i r e t o n t t f c h m i l o o t h e r t a n t s w i t h t h e s k i n L e s o n m o t o f t n o n e p o s e d p r i t s a n d m y b a s y m m t i c a l (f d t o u n t n a l g e n t s) L e s i o n a g g r v a t d b y x p o s u r t o t h i r r i t a n t a n d t h s h o l d b a v o i d e d P a t h t i s m y b o f v a l u e i n d g n s e c o o b a t i o n o f l i n i a l i m p r e o n s

Diagn s

S u v y t h p t i t s v i n m n t a n d s t u d y h t o t l a t i v i t i e s t d t e r m i n i r r t a n t

A S c h f a l i s t y o f r e n t e x p r t n e w h e m i c a l d u g s s o a p c o s m e t r o t h o n t t i i t a n t s T h e l c a t i o f t h l i a l m y b e f v a l u e i n i d n t i f y i n g t h i t a n t e g l p (r u n e o h m p o o) f (s o p s h v i n g m t i l o m t i) e c k (j e w l r y c l o t h i n g) t r u n k (c l t h i n g) u p p e x t m i t i (p a o m t i p l a n t t o x i n i n d u s t a l h m a l) a n d l o w e x t r e m i t i (t o c k i n g s h o e s h o e d y e s)

COMMON DISORDERS OF THE SKIN

PRURITUS (Itching) (code No 143)

Treatment

- A Specific Measures** Remember that localized (as well as generalized) pruritus may result from systemic causes
- Remove or treat specific causes whenever possible
- 1 Skin infestations (e.g. scabies, pinworms, pediculosis)
 - 2 Skin infections (e.g. fungal and bacterial infections)
 - 3 Skin inflammations (non infectious) (e.g. lichen planus, eczema, urticaria)
 - 4 Altered sweat secretion (e.g. hyperhidrosis, anhidrosis)
 - 5 Allergic reactions (e.g. food, drug, clothing, serum, etc.)
 - 6 Senile dermatosis (e.g. senile skin atrophy)
 - 7 Metabolic disease (e.g. diabetes, hyperthyroidism, goiter)
 - 8 Uremia
 - 9 Jaundice
 - 10 Opiate intoxication (e.g. morphinism)
 - 11 Blood and neoplastic diseases (e.g. leukemia, lymphoma)
 - 12 Psychogenic factors (e.g. anxiety states)

B Local Measures

- 1 Shave lotion, emulsions and ointments incorporating the volatile analgesics and antipruritics listed in tables of pages 100 and 107 may be of value in relieving itching
- 2 Relieve excessive dryness or moistness of skin
 - a If skin is too dry, softening agents may afford relief e.g. use water ointment (R 31 page 103). An excellent principle for dry skin is to wet it first in a bath (to hydrate the keratin) then apply petrolatum to the wet skin to trap the moisture
 - b If skin is too moist, drying agents may afford relief e.g. wet dressing, socks (R 18 page 98-99), hker lotions (R 14-16 page 100) and powders (R 9-12 page 99) (especially if pruritus is acute)

Tar baths. Generalized pruritus may be effectively controlled by lukewarm baths 15 minute bid or tid. After bathing the skin should be blotted not rubbed.

 - (1) Starbath and sodbath 1-3 cups tarh and 1 up soda um bicarbonate dissolved thoroughly in tubful (50 gallons) of lukewarm water (Soda may be omitted)
 - (2) Tar baths D: olv 50 100 Coal Tar Solution U S P in one tubful (50 gallons) of warm water (Watch for sensitivity)

CAUTION Avoid excessive drying of skin by overbathing prolonged bathing periods and exposure to drafts after bathing

C General Measures

- 1 Diet
 - a Food should be simple. Avoid rich and spicy foods
 - b Test diets or elimination diets should be used in cases of suspected food allergies (see page 56)
- 2 Psychotherapy If pruritus is primarily a manifestation of an anxiety state, obsessive-compulsive or psychtic

ERYTHEMA NODOSUM (Due to Infection code No 114 1x0)

A tender nodular erythematous dermatosis occurring most commonly on the extensor surfaces of the legs and (less often) for arms. It is usually caused by toxins of infections and occasionally by drugs. The disease occurs most commonly in the spring or fall and usually runs a course of 2-6 weeks or longer.

Treatment

A General Measures

1. Eliminate or treat the specific cause.
Infections: Almost all infections (coccal, tuberculous, mycotic or viral) are capable of causing erythema nodosum. For treatment see specific diseases.
2. Exogenous toxins: e.g. drugs or chemicals.
3. Rest: Hospitalization may be advisable.
4. Focal infections: May be corrected although this does not appear to influence the course of the disease.

B Local Measures: Usually unnecessary but if lesions are troublesome or complicated treat according to stage and type of dermatitis (see pages 93-96-97).

C Topical drug: 250 mg q.i.d. for several days have been shown to be effective in some cases (empirical data).

D Steroid therapy may be used if not contraindicated (tuberculosis must be ruled out). Gv. Repository Corticotropin [injection] 10 USP 20-40 units I.M. daily or very high dose for 2 weeks or a course of the corticosteroids (generally in large doses).

ERYTHEMA MULTIFORME

(Infection code No 11x 190) (Poison code No 11x 3x7)

An acute inflammatory polymorphic kind of multiple and sometimes deep. There is often a history of drug poisoning or of a recent infection. The skin lesions found most frequently on the distal parts of the hands and feet and on the face. The lesions are usually limited although they frequently recur.

Treatment

A Drug Measures

1. Eliminate causative factors.
Chronic systemic infections (e.g. tuberculosis).
2. Focal infection.
Satisfying drug.
3. Penicillin: 1-1,500,000-3,000,000 units I.M. as indicated by the extent and duration of the illness when a satisfactory response is not obtained (CAUTION).
4. Oxytetracycline: 250 mg q.i.d. for 1 day may be useful.
5. Corticosteroids may be used as for erythema nodosum.

B General Measures: Bed rest and good nursing are very important.

C Local Measures: Treatment and type of dermatitis (see pages 93-96-97). The following principles should be observed:

- B Use protective isolation in certain selected cases cautious re exposure may help to establish the irritant
- C Patch tests may be of value but false positive and false negative reactions may occur Dermatitis produced by such tests should resemble the clinical dermatitis In the event of a positive reaction a control test should be done on a normal individual

Treatment

A Definitive Measure

- 1 Prevent re exposure to irritant
 - a Avoid soaps and detergents
 - b Cosmetics Change to so called non allergic cosmetics or eliminate cosmetics entirely
 - Occupational irritants
 - (1) Protective rubber gloves may be used but are seldom indicated In such cases an inner cotton glove must be used
 - (2) Protective creams (barrier creams) may be tried but are of limited use
 - (3) Change of occupation or duties to those not involving use of irritants may be necessary
 - d Plant irritants (especially Rhus species e.g. poison ivy)
 - (1) Destruction of plants by manual removal or by chemical means (2,4-D or d-chlorophenoxyacetic acid) near dwelling and areas frequented by people
 - (2) Avoidance of Rhus infested areas
- 2 Prompt and thorough removal of irritant by prolonged washing or by removal with solvent or other chemical agents may be effective if applied very shortly after exposure In the case of Rhus toxin thorough washing with soap and water must be done within a few minutes if it is to be of any value

B Local Measures Treatment and type of dermatitis (See pages 93-96-97)

- 1 Acute weeping dermatitis
 - a Do not submerge with soap and water
 - b Apply soothing solutions (see table on page 98) If the patient becomes generalized use the soothing starch and barborat antipruritic bath mentioned on page 66
 - Shaklotins (§ 14-16 page 100) may be indicated instead of wet dressings or in interstices between wet dressings especially in interdigital areas or when oozing is not marked Lesions on the extremities particularly may be bandaged with wet dressings
 - Hydrocortisone and fluid corticosteroid preparations (35 different preparations of hydrocortisone and other materials) lotions cream ointments topical pigments and dimethylmethylpyrrolidone

SABAUTE dermatitis (subdiagnosis) Use shake lotion

- 3 Chronic dermatitis (dry and lichenified) Treat with hydrophilic glycol ointments or cream Tars or phenol preparations if in a stage of the dermatitis

C General Measures Repository Corticosteroid Injection U.S.P. (corticosteroid) or on of the cortisone may be tried (CAUTION) and repeated daily (see page 98)

- 2 Cortisone in lotion cream or ointment for m applied sparingly twice daily may be very helpful (see page 424)
 - 3 Treat the clinical type and stage of the dermatitis
 - a Acute weeping lesions Use solutions listed in table on page 98 as soothing or soothing soaks baths or wet dressings in the daytime for 30 minutes tid or qid Shake lotions (§ 14 15 page 100) may be employed at night when wet dressings are not desirable Lesions of exfoliative dermatitis may be damaged for protection at night Powders (§ 9 11 12 page 99) may be used in intertriginous areas when ooiling is not marked
 - b Subcutaneous or abscessing lesions may be treated with saline lotions which may incorporate mild antipruritic or mild stimulating agents (see page 107) Shake lotions are usually preferred for widepread lesions Ointments (see page 104) containing mild tar may be used (see table on page 106)
- Chronic dry lichenoid lesions are best treated with ointments creams and pastes (see pages 102 3) employing lubricating keratolytic antipruritic and mild kerosene topical agents mentioned in the table on page 106 7
- Indicated The treatment is perhaps the most popular therapeutic agents in chronic eczema (25% ointment in ointment cream and pastes) Iodochlorhydroquinone U S P (Vioform®) 3% o Chloroquinolone N N D (Stessa®) ointment to cream may be indicated especially if the antipruritic is not sufficient
- 4 X-ray therapy (by specialist) may be used effectively if only temporarily in many stages

DERMATITIS MEDICAMENTOSA (Drug Rash) (code No 110 3)

An acute or chronic inflammatory reaction on which is caused by a wide variety of drugs and which causes a wide variety of skin lesions in susceptible individuals The reaction may be immunologic or idiosyncratic (idiosyncratic) and may or may not be associated with constitutional disturbance (fever headache etc) Improvement following withdrawal and elimination of the suspected drug usually takes a few days but may take longer A rule it is not advisable to attempt diagnostic provocation of an allergic reaction by re-exposure to the drug Skin tests are of little value

Treatment

A Symptomatic Measures

- 1 Stop all drugs if possible
- 2 Have elimination of drug by increasing fluid intake
- 3 Give symptomatic relief agents
 - a Diminution of U S P (BAL®) may be tried in severe to heavy metal (arsenic mercury gold etc) (see page 536) Edthamil Chloride N N D (Venat®) may be worth trying for lead poisoning (see page 541)
 - b Sodium chloride 5 10 Gm (75 150 g) daily orally may hasten elimination of bromides and iodides and lead to the drug (see pages 538 and 540)

70 Eczema

- 1 Acute lesions Employ simple wet dressings and soaks or soothing lotions For treatment of buccal lesions see page 261
- 2 Subacute lesions Soothing lotions

Prophylaxis

Avoid all unnecessary medication in susceptible individuals
1. Patients with a previous history of erythema multiforme

ECZEMA (code No 111.390)(and Eczematoid Dermatitis)

A large group of non specific acute or chronic superficial inflammatory skin reactions which occur as a result of exposure to chemical physical or unknown irritants or as a result of allergens. Irritants may be external (e.g. contact dermatitis) or internal (e.g. dermatitis medicamentosa). There may be a history of allergic tendencies (atopic eczema) and blood eosinophilia may be found. The term eczematoid dermatitis is used for eczema like reactions of undetermined origin. The lesions frequently are usually pruritic. Acute lesions are usually erythematous vesicular or exudative. Chronic lesions are usually thickened squamative or lichenified.

Treatment

A Specific Measures

- 1 Elimination of inciting agents (see above) is in a sense the only specific measure. A careful history trial and error elimination and exposure techniques may be of value in incriminating possible offending agents. Skin tests are often valueless. Desensitization is of no value. Sensitivities are usually multiple.
- 2 Diet Should be adequate and well balanced. There is no evidence to suggest that standardized or routine dietary restrictions are of value especially in adults. Trial diets or elimination diets may be of value in determining food allergens in individual cases when an urticarial component is present. Food diaries may be kept by patients with chronic eczema to determine possibility of food allergy. Reported common food offenders are wheat milk eggs pork fish shellfish tomatoes strawberries and chocolate.
- 3 Psychotherapy An attempt may be made to determine and correct existing emotional disturbances but this is of no practical value.
- 4 Remove definite foci of infection but avoid routine polysurgery.

B General Measures Corticotropin (ACTH) or the corticosteroids may provide symptomatic improvement in severe or fulminant eczema (see page 424).

C Local Treatment

- 1 Avoid all unnecessary local irritations to the skin such as may occur from excessive bathing, or as a result of exposure to irritating drugs chemicals greases and soaps. Soapless detergents are not advisable. Clear up skin infections promptly (particularly those with pus) by appropriate measures (see pages 84-85).

- 4 Topic l ti infecti e drugs [g 1% squ s n omyc
o yt tr cy li e hl tetra y line hloramphe i ol
(CAUTION) ythromycin o polymyxin B o (ments)
ho ld be us d when n e sary (see p ges 84 86 a d 107)

P ophyla

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DERMATITIS ACTINICA (code No 110-451) (Erythema Solare or Sunburn)

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T t m i

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83) F st col g d sooth g wet d ss i gs (s pag 98)
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q Pho ph t U S F 0 25 Gm (3 3/4 g) b i d now
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daily o Gold Sodium Thio ulphate N F 50 mg (3/4 gr)
I V o ce weekly (CAUTION) may be t ed

P ophyl

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l m i a y o d t i o g by g a d d p s is d i b l
B P o t i Ag t Apply to ki b f pou e t d t i o n
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2 C bolat d (phen liz d) V lin ® s a good s n
3 Menthyl a thra ilat (5%) a d 5% t t ium d o de e m
4 D g lloyl tri leat cr m (N o A fl®)

LICHEN PLANUS (code No 110 965)

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7 Exfoliative Dermatitis

B General Measures

- 1 Discontinue all unnecessary medication when feasible for as long a period as possible
- 2 Treat systemic manifestations as they arise e.g. anemia, icterus, purpur, etc.
- 3 Antihistaminics may be of value in reactions of urticarial and angioneurotic character (see page 66)

C Local Measures Treat the varieties and stages of dermatitis according to the major dermatitis which is simulated

- 1 Eczematoid (see page 70)
- 2 Acneiform (see page 77)
- 3 Puritic (see page 68)
- 4 Pyoderma (see page 84)
- 5 Urticarial (see page 78)
- 6 Bullous (see page 88)
- 7 Lichenoid (see page 73)
- 8 Exfoliative (see below)

EXFOLIATIVE DERMATITIS (code No 110 966)

A serious cutaneous reaction often due to sensitization to certain drug (e.g. arsenic and gold) and also commonly caused by lymphoblastoma. It is characterized by itching, weeping erythematous patches which rapidly coalesce and spread to become generalized. Finally a desquamation or exfoliation of large areas of skin occurs. There is an associated severe constitutional reaction with fever and other systemic symptoms. The disease runs a course of weeks to months and is attended with a high mortality rate.

Treatment This is a medical emergency

A Specific Measures

- 1 Stop all drug if possible
- 2 Hasten elimination of offending drug by all means e.g. by inducing fluid intake
- 3 Diminution of U.S.P. (BAL®) may lessen the severity or duration of reaction due to arsenic or gold (see page 536)
- 4 Corticotropin (ACTH) U.S.P. 20-40 units I.V. or Cortisone A.T.A. U.S.P. 50-100 mg I.V. orally if indicated

B General Measures

- 1 Hospitalize patient when possible. Use talc or bed sheets
- 2 Keep room warm, comfortable. Avoid drafts
- 3 Institute supportive treatment as follows: plasma
- 4 Avoid all unnecessary medication
- 5 Corticotropin (ACTH) or cortisone may produce symptomatic improvement in severe or fulminant exfoliative dermatitis (see page 424)
- 6 Secondary infections. Penicillin or other antibiotic drugs should be given when there is evidence of bacterial infection (see pages 84-86 for dosage schedules). Pyoderma is the most severe complication of exfoliative dermatitis

C Local Measures

- 1 Observe careful skin hygiene
- 2 Avoid irritating local applications
- 3 Treat skin as for acute extensive dermatitis
 - a First: Wet dressings, soothing baths (see page 66), powders (see page 89) and hake lotions (see page 100)
 - b Later: Soothing oily lotions (see page 100) and ointments (see pages 102-103)

- 4 Topical antifungal drugs [eg 1% aqueous nystatin ointment] or chlorotetracycline or rifampicin
(CAUTION) erythromycin or polymyxin B ointment]
should be used when necessary (see pgs 84 86 and 107)

Phyl

Phyl is a red to brown drug should be withheld if
fully formed development of skin lesions of all types. The drug
should be withheld until the state of the skin reaction is determined.
Definiteness of reaction may be obtained by an absolute contraindication
to further drug administration.

DERMATITIS ACTINICA (code No 110-451) (Erythema Solare or Sunburn)

A acute inflammatory skin condition following exposure to a large
dose of ultraviolet radiation. It may vary from simple erythema
to severe exfoliation and may be associated with systemic manifestations.
Some individuals are abnormally light sensitive.

Treatment

- A Symptomatic Management. Treatment of symptoms by
topical application of emollients. Cool compresses in buccal
dilatation to relieve the symptoms of dryness.
B Local Measures. Treatment of erythema (see pgs
93) First cool compresses with water (see pgs 98)
follow with lotions (see pgs 100). Continue treatment until
healing of the lesions is effected.
C For polymorphous dermatitis. Light sensitive eruptions. Chloro-
quine phosphate USP 0.25 Gm (3 3/4 g) bid now
apparently the best treatment for holoferon. For severe
intense cases. Chloroquine 500 mg bid. 500 mg bid. 500 mg bid.
daily of Gold Standard Thio sulphate NF 50 mg (3/4 g)
1 V o c e weekly (CAUTION) may be tested

Phyl

- A Individual with erythema sensitive skin should avoid exposure
to prolonged exposure to the sun or ultraviolet radiation. P
liminary conditioning by gradual exposure is advisable.
B Potent Agent. Apply to skin before exposure to radiation.
1 Potassium permanganate Acid NF 10% hydrophilic ointment
C Bland (phenolic) Vaseline ointment
3 Methyl salicylate (5%) and 5% tincture of menthol
4 D Glycerol ointment (N o A f l)

LICHEN PLANUS (code No 110 965)

A chronic inflammatory skin disease of unknown cause
characterized by small flat topped violaceous papules which
are angular in shape (usually quadrilateral) and of varying sizes.
They commonly occur on the surfaces of the arms and
in the subcutaneous tissue of the back and the genitalia.
There may be associated bullous lesions. Readily pigmentation and
atrophy may be associated with the lesions. Lesions of the
skin may be simulated by drug reaction (benzothiazine)

74 Psoriasis

Treatment.

A General Measures

- 1 Phenobarbital U S P 15-30 mg ($\frac{1}{4}$ - $\frac{1}{2}$ g) b i d q i d
- 2 Psychotherapy Patients are often high strung or tense and nervous. Episodes of dermatitis may follow emotional crises. Measures should be directed at relieving anxiety.
- 3 Chloroquine Phosphate U S P 0.25 Gm ($3\frac{3}{4}$ g) b i d orally for one month is worthy of trial. If chloroquine is not tolerated, hydroxychloroquine sulfate (Plaquenil®) 0.2 Gm (3 gr) b i d orally may be tried.

B Local Measures

- 1 Use shake lotion containing tar (§ 17 page 100)
- 2 X ray may be used only in severe cases which have proved refractory to other forms of treatment. Treatment by x ray must be reserved for the specialist.

PSORIASIS (code No 111 961)

An acute or chronic inflammatory skin disease of genetic etiology which is characterized by macular and papulopustular lesions of varying sizes and configuration (usually with well defined borders). The lesions have dry silvery scales and bleeding occurs when these scales are removed. Pruritus is rare except in flexures or in acute eruptive cases. The lesions occur on the extensor surfaces of the extremities and on the trunk, nails and scalp. There is sometimes an associated disfiguring arthritis but a constitutional factor. Stippling of the nails may be pathognomonic.

Treatment

A General Measures

- 1 Climate Warm climates seem to exert favorable effect
- 2 Non-specific internal medication of little value with the exception of arsenic which is hazardous in a few of the patients at the rate of the lesions and the daily effect of excretion of arsenic (k ratio is pathologic)
- a Cyanocobalamin U S P (Vitamin B₁₂) 1000 mcg I M 2-3 times a week is said to be of little value
- b Arsenic Fowler's solution (Potassium Arsenite Solution N F) has been recommended in doses of 3-15 drops twice daily in patients with subacute or chronic lesions. Dosage duration of administration on indication and evaluation of the drug are of doubtful value. It may be given in repeated courses if desired but the results should be waited for 2-3 months (see page 239)
- 3 Psychotherapy Response is important since the symptoms are apt to be discounted due to chronicity of the disease. An attempt should be made to relieve existing anxieties.

B Local Measures

- 1 Acute psoriasis (Avoid irritating or stimulating drugs)
 - a Begin with a shake lotion (§ 14-15 page 100) or bluish ointments (see page 102) containing 5% detegent solution of coal tar
 - b As lesions become less acute

mil

ke at pla t c ag nt (s p g 106) to lotions (see pag 100) and hyd oph l ointm ts (see page 103) W tch p t t ar f lly

2 Seba tep o las

- a Give wa m baths da ly ac b b ng the ski lesions tho ghly with b ush o p a d wat r
- b Apply crea g o centratio s of k r topl tie or st mu lating gents (e p ges 106 a d 107) i o po at d in l tie s (see page 100) a d hyd phili ointme ts (see page 103)
- Sol r o ultr iolet i r d ations m y be ppl d ing d ally in si g dos

3 Chro psoriasis

- Amm iated Mer u y Ointment U S P (57) l cally b i d
- b A th al N F 1/47 ointment lo ally o c a day (d y)
- c Comb d ult avi let i r d at on d t r g men (m d fied f m G kerm) (Daily n ed d) Smear 2 57 lta ointm t (s p g 71) th kly on ki d l e f 12 24 h u s Wip off o tm nt with miner l il l av g l ght t l Foil w with da ly gr d d s b r yth m do es f ltr let l ght as t l ted

PITYRIASIS ROSEA (code No 111 962)

A c mmon mild ac t inflammato y skin d s ase of unkn wn ti l g y wh h is h a te i ed by p p lo quam o erupt n on the trunk m s d th ghs and wh h o curs m r f equ ntly w th p ing and f ll The pap le are pink and oval w th ac ling bo d and p l e te th y a e typically a ang d w th th i lo g al gth le v g lines f the ki A ngl h rald p t h m y p ec d multipl l ion by a pe i d of sev l d y The l sion may or m y not be p urit The d seas usually l ts 6 we k w th without tr tm t

T sim t

A G l M u None

B Loc l M

- 1 Acute it t dle i s a uncommon l i p es nt tre t a f r a te derm tit with wet d eal g s (see p ges 98 and 99) o w th h k l tion (B 14 17 pag 100 101)
- 2 Cal Tar Sol i on U S P 57 n st ch lot b i d
- 3 Ultr iolet light is h l p l
- 4 Pre it s S lo l stip iti meas r on pag 86

SEBORRHEIC DERMATITIS (code No 111 190)

An a ut ch o ic p pul q mo d m titis ften as ocl t d w th c siv o lin of th kin and co ing in th so c l l d b o a as f the b dy (elp midporti of f at nal gion and int re p l region) Th le so ppe (1) a y il w h gr sy les o (2) an ut or h o i c m tons d rmatitis in e s of seba ou gland e c t ti and in intert ignou r a e tly a p ur ti

TreatmentA General Measures

- 1 Diet Well balanced adequate diet avoiding excess sweets spices hot drinks and alcoholic beverages
- 2 Regular working hours recreation and sleep
- 3 Simple cleanliness
- 4 Remove aggravating systemic factors (infectious overwork emotional stress constipation and dietary abnormalities)

B Local Measures Treat type and stage of dermatitis

- 1 Acute subacute or chronic eczematous lesions Treat generally as for dermatitis or eczema (see page 6)
- 2 Seborrhea of scalp
 - a Selsur® suspension (selenium sulfide) following weekly shampoo Foster® cream (containing soapless cleansers wetting agents hexachlorophene sulfur and salicylic acid) may be used as a weekly shampoo for oily seborrhea
 - b Sebizon® lotion (sodium sulfacetamide) once daily
 - c Mild coal tar scalp lotion (§ 21 page 101) may be used
- 3 Seborrhea of non hairy areas Mild stimulating lotions (§ 17 page 103 or 20 page 101) may be used Ointment (§ 36 page 104) or 3-5% sulfur in hydrophilic ointment (see page 103) may be used (The addition of 1% salicylic acid aids in removing scales)
- 4 Seborrhea of intertriginous areas Avoid greasy ointments Astringent wet dressings (§ 16 page 98) followed by 5% ammoniated mercury in hydrophilic ointment (see page 103) may be used

EXTERNAL OTITIS (code No x75 100)

This may be considered a variant of seborrheic dermatitis and at times may become an infectious eczematoid dermatitis. An interference with ceruminous secretion leads to inflammation of the canal wall and predisposition to secondary bacterial infection usually with *Pseudomonas aeruginosa* (*Bacillus pyocyaneus*). Fungi are rarely if ever causative.

TreatmentA General Measures

- 1 Penicillin 300 000 units once or twice daily I.M. for accompanying fever and erysipelatous changes (CAUTION)
- 2 Phenobarbital U.S.P. 15-30 mg (1/4-1/2 g) b.i.d. q.i.d.

B Local Measures

- 1 Aural lesions Coal wet dressings
- 2 To remove cellular debris if present Glycerin of hydrogen peroxide with carbamide as ear drops b.i.d.
- 3 Iodochlorhydroxyquin U.S.P. (Vioform®) 3% cream b.i.d.
- 4 Hydrocortisone Acetate Ointment U.S.P. 1-12% locally
- 5 X-ray therapy in refractory cases (must be given only by a trained specialist)
- 6 Polymyxin B-bacitracin ointment (Polysporin®) oxytetracycline chlortetracycline neomycin or erythromycin ointments (see page 514)
- 7 Corisporin® otic suspension (hydrocortisone neomycin polymyxin B) may be beneficial when eardrum perforations fail

ACNE VULGARIS (code No 151 7x0)

A mmo infl mmato y kind se of gen ti origin p o
voked by androg s in the mal and p ogest e in the f mal It
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T eatm t

- A G l Meas El minat all unn s ry m dicat n
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 - 2 A id o p t on l expos e to msn ral oils and g ses
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for o w k (pr m st ually) ch month
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1 2 mg (1/50 g) d ly o
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mg (1/40 g) d ly
d Eth nyl Estr d ol U S P 0 01 0 05 mg /c in 70%
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ea h d y for 3 months m y b tr d b t has l mlt d valu
 - 5 C r t y t m c d g me t l d g tion nat p tio
ml trit f ti a m dem t l d turb c a
 - 6 V Aut ge o a d sto k v ies d ther f reign
p ot n t g sh e b en employ d w th qui o l r lt
 - 7 T t cycl e U S P 250 mg d ly e y th day fo
w ks o mo ths v y helpf lf ome e fa e

B L l M es

- 1 Local lean ing of sk n and calp
a O d y so p f leans g
b A id g easy la g r ms doth r m ti a
c Shampoo alp l 2 t me w k (R 48 p g 105)
- 2 E tr tio and d in ge flo al les o s in s l t d ca s
a Ext ct bl ckhe d w th om do xtr t rafte oft ning
f e w th h t w ter compr s fo 1/2 l ho r
b In and d in fl ciuant y tic lesi n s w th small sh rp
lp l H t mp s l p hov t i d f vor d in g
- 3 K t pl st d k t lytic ag nt
a Hot wat o b ric a id c mp s s (not teaming) m y
b us d to p od hype mi and desq am t on of l i n
b K ratolyt lot ons A ne l tion (s lf r lotio) (R 19
p g 101) o alf inoll ion (R 20 p g 101)
may b tr d th ya appli d loc lly to the kin t bed
time a d wa h d off i th m ni g
c Kerat lyt intm t and p t B gun w th w k p epa
t ons and bu ld up as t lerated Apply t b d tim and

remove in the morning

(1) Sulfur 2-10% in hydrophilic ointment (see page 103)

(2) Sulfur and kaolin paste (p. 39 page 104)

(3) Quinolol[®] ointment or Vioform[®] ointment (see page 107)

4 Irradiation

a Simple exposure to sunlight in graded doses is often beneficial

b Ultraviolet ray May be used as an adjunct to other treatment or to remove scars Use suberythemal doses in graded intervals up to point of mild erythema and scaling

c X rays This is very effective technique should be reserved for only the more severe cases and only after other more conservative measures fail X-ray therapy must be reserved for the specialist

URTICARIA (Hives) (code No 11x 390) and ANGIONEUROTIC EDEMA (Giant Hives) (code No 11x 580)

An acute or chronic inflammatory skin reaction of allergic origin manifested by multiple markedly pruritic wheal reactions of varying sizes with no characteristic localization and at times involving the mucous membranes. Acute attacks are usually self-limited from a few minutes to a few weeks but have a tendency to recur. In extreme cases a laryngeal edema may cause a respiratory obstruction and death. Skin infection, contact dermatitis, toxic erythemas, etc. may be other causes of urticarial reactions.

Treatment

A General Measures

1 Purgation Initial purgation to remove possible antigenic material has been recommended in acute cases. Castor Oil U.S.P. 15-30 cc (1/2-1 oz) may be given. Stool may be examined for parasites.

2 Diet During the cutaneous phase the diet should be simple and free of such common offenders as wheat, milk, eggs, pork, fish, shellfish, tomatoes, strawberries, and chocolate. Past history of food allergies should be ascertained and elimination diets may be helpful in determining the offending food. The patient should maintain a record of diet and symptoms. The patient should be advised that known food offenders are:

3 Avoid use of sympathomimetic. Suspect lidocaine (even epinephrine, ephedrine, histamine, cocaine, etc.)

4 Drug

a Antihistaminic drugs often give prompt and sustained symptomatic relief. If penicillin which is being given as a lifesaving measure produces an urticarial reaction it is sometimes possible to continue the drug by simultaneous administration of an antihistaminic drug. This must only be done with great care. Examples of commonly used antihistaminic drugs are given on page 45.

b Epinephrine Injection U.S.P. 0.3-1.0 (0.15-0.45) of 1:1000 solution but feel only when (1) Laryngeal edema is present

() U t c i a i i n t s e

(3) A t i h a t a m i d r u g s h a e f a i l d t o g v e e l f

c Ephedrine S lfate U S P 25 mg ($\frac{3}{8}$ gr) or lly q i d

d Ephedr e edati e m t r f o l h p y o p r p h y l a x

(o e c a p u l q i d)

(1) Ephedrine Sulfate U S P 25 mg ($\frac{3}{8}$ gr) and

P ntob r b i t l Sod m U S P 25 mg ($\frac{3}{8}$ g)

() Eph drine S lfat U S P 25 mg ($\frac{3}{8}$ gr) and

Phe barbital U S P 15 mg ($\frac{1}{4}$ gr)

5 Co tic tropin (ACTH) or the co t i s o e s m y p o v i d e s p e c t i l r i m p r o v e m e n t s e r e o f l i m i n a t a n g i o n r o t c e d e m (s e p a g e 423) T h e s e d r u g s s h o u l d b e e d o n l y i f i t i s a p p a r t h t h t h p a t t w i l l n t r e s p o d t o m o e c o n s a t v e m s s

6 M i s c l l e o s m e a s r e h a b n r o m m e d d f o r t h h r o f o m o f t h e d s e a e b t t h e i r v a l u i s q u t i o e d a D i l t d H y d o c h l o A c d N F 15 20 d p a t i d a a n d d u n g m l s B r a h t e t h a f t r m e l s w i t h o d m b a r b o a l e

b C l c i m G l u c o n a t e U S P 1 G m (15 g r) t i d

o r a l l y p e O t h c a l i u m a l t s m a y a l s o b u s d

B L o a l M e a s e A n t i p r i t s a f r e q u e n t l y o f b n f i t

1 S o o t h i n g t i p u r t c b t h s (s e e p g 66)

2 S o o t h i g a t i p i t t i c l o t i o (s e e p a g e 100)

Prophyl is

A E l i m t d v o d e p o u t o c a t v e f c t o

1 S e t g d g A l m o t a l l d g a r e a p a b l o f p r o d u c i g a n t i i a l r a t o O p t b r b i t u r t e s s l i y l t a p i l l s u l f o m d e s b r o m i d o d d a t i h i t a m c c o t i c o t o p i n (A C T H) t

2 S e t i z i n g f o o d A y f o o d m a y p r d u c e a u r t c a i a l e p o s e s c e p t i b l i d i v d a l s a n d s h o u l d b e c o n s d e r d i n o b c u e a (p t i l r i y h c c a e s) f u t c

3 A g g a t i g p h y i l f t o g c s h e a t n d o l d k i n a n d m u c o s m e m b i r t t e

4 A g g a v a t i n g s y s t e m i f c t o s g c h r o n i i n f e c t i o n s f o i o f i f c t o n s p i t i i f e t t i o a d b l o o d y r a s

B R l i f f P y c h i D i r b e l u s c p t b l e d i v i d u a l m o t o a l t e a d t m a y p r p t a t t h i l o n s

INTERTRIGO (code No 111 437)

E y t h e m d e t o c h f i g o f t h s k i n

T i m e t

T e t s t i n a c r u r i s (s e e p g e 89) b u t d o o t e f g a d l g e t

MILIARIA (Heat Rash) (code No 153-445)

An acute dermatitis characterized by small erythematous burning and often pruritic papules, vesicles and pustules which occur most commonly on the upper extremities, trunk and intertriginous areas. It is caused by exposure to a hot moist environment.

Treatment

A General Measures

1. Provide optimal working conditions when possible i.e. controlled temperature, ventilation and humidity.
2. Avoid overbathing and use of strong irritating soaps.

B Local Measures

1. Antipruritic cooling lotion apply b i d to q i d

Rx	Menthol	1 0	gr xv
	Phenol	2 0	ss
	Glycerin	15 0	3 v
	Alcohol 35% q s ad	240 0	3 viii
2. Drying shake lotion (Rx 14 with 1% phenol or Rx 15 page 100)
3. Sulfur resorcinol lotion (for seborrheic skin) (Rx 20 page 101)
4. Antipruritic powders or other dusting powders (see page 99)
5. Treat secondary infection (pyoderma superficial) with potassium permanganate soaks, compresses or baths (see page 98). Ammoniated mercury 2.5% in a hydrophilic ointment (see page 103) may be employed distally.
6. Tannic Acid N.F. 10% in 70% alcohol locally b i d.

Prophylaxis

- A. Trough skin. Guarded exposure (or sedalys) to light or ultraviolet light may be effective in which will be subjected to hot moist atmosphere.
- B. A cold position to avoid atmospheric conditions of extremely high humidity.

ANO GENITAL PRURITUS

(Ani code No 143 573) (Vulvae code No 771 570)

Diagnosis

- A. Consider the role of systemic lesions of pruritus, anxiety states, diabetes, trichomonas infection and nitelliparasitosis.
- B. Rule out all obvious local pathological conditions of the anus and rectum, bowel irregularities, colitis, tanus, etc.

Treatment (See also Pruritus page 68)

A General Measures

1. Diet. Avoid hot spicy foods (e.g. hot peppers, chili) and drugs which are irritant to the anal mucosa.
2. Treat tip of the penis (see page 254).
3. Provide necessary psychological treatment as indicated.
4. Instill the patient to use very soft or moistened tissue or cloth after a bowel movement and to clean thoroughly. Women should apply antipruritic ointment after urination.
5. Instill the patient regaining harmful and pruritus inducing effects of scratching.

B L o i M u e s

- 1 Phenol ted Cal mt e Lot on U S P ppl d locally
- 2 S t b t h s b i d if the a s is a ut ly i flam d d o o i g
us g S l v N t at U S P 1 10 000 1 200 (0 01 0 57)
Pot ium P manga ate U S P 1 10 000 (0 017) o
Al m n m Sub cetate Sol t n U S P 1 0 (5%)
- 3 U der l th g h l d b cha g d d a ly
- 4 C t o l x s a p r s p r t n by u e of d ying p wd s such
as talc (s page 99)
- 5 P a i t f s s u d o r u l t d as w th S l v N t ate U S P
5 107
- 6 Hydroco t one A et te O r t m nt U S P 1 2 1/27 locally
b d
- 7 X r y t h e r p y may b u s d if o t h m u r e s f i l This
sh uld b r e s e e d f r the s p i a l t

P o p h y l i a x i s

- A T e t a l l p s s i b l y s t m i c o r l o c a l u s e
B I n s t r u t t h p t i e t i n p r o p a n o g e n t l h y g n e

**CALLOSITIES (code No 112-430) and
CORN (of feet or toes code No 148-433)**

T r t m t

- A C o r t m e h n i c i b m l y c i s w h i t h u s f r i c t a n d e
ult in th ho y o v growths
- 1 Sho s m u t b p r o p l y f t t e d
 - 2 O t h o p d i c d f m i t t i m u s t b e t r e d a n d c o r r e c t d
- B R m o v C l i t By
- 1 P u n g o f l l a f t w a r m w t e k
 - 2 K e t l y s i b y u s o f h m l a g n t s

a B S l i c y l c a i d	4 0 3
A t o n	4 0 3
C o l l o d o n q s a d	15 0 3

S g A p p l y l l y t o a l l u v r y n i g h t a n d o e w t h a
t i p o f a d h s i R m o a d h s i v i n t h m u n g R e
p e t u n t l c o r n o r l l u i m o v e d

b C o m m c i a l s a l i y l a c i d c n p l s t m a y b u d
 - 3 A m e t a t r l i e t h e r b a r 1/2 i h w d a n d 1/4 i n h h i g h m a y
b p l e d o n t h o u t i d e o f t h h o e j t b h i n d t h e w l i g h t
b a r i n g u f e o f t h s o l

DRY SKIN (Congenital Senile or Environmental)

T r t m tA G l l i s t r t i to P t i e t

- 1 A v o d e x e s s v e b t h i n g a n d u n o o s p A v o i d u n d u q y
i n g i r t a t i n g o k t l y t i c m e d a m e n t a v o i d o l d o
d y e v i r o n m n t
- 2 A p p l y m p l g s i b l l y t o t h e s k i n w h i l i t i w e t
c o o n t b u t t e g t b l c o o k i g f a t s H y d o u W o o l F a t
U S P (1 a l l) L q u d P t o l a t u m U S P (m i a l l)
P t r o l t m U S P o r i m p l o i t m t s (p g 102 103)

- 3 Soapless detergents may be used when bathing but they may do more harm than good

B General Measures

- 1 Treat complicating dermatoses (e.g. scalp eczema and pyoderma) by appropriate measures (see pages 71 and 84)
- 2 Vitamin A in high doses (50 000-100 000 units daily) has been recommended but results are questionable

HERPES SIMPLEX (Cold or Fever Sore) (code No 13 166)

An acute viral infection apparently precipitated by various causes such as fever infection allergy ultraviolet radiation and psychic trauma. The small grouped vesicles can occur anywhere but are most frequent on the skin and mucous membranes of the face nose mouth throat and genitalia. Regional lymph nodes may be involved. Attacks are usually self limited but are often recurrent.

Treatment

For persistent or severe recurrent herpes

A General Treatment

- 1 Eliminate precipitating agents when possible
- 2 Routine smallpox vaccination twice yearly intervals for 6-8 weeks. Equivocal results

B Local Measures

- 1 Dust vesicles twice daily with bismuth formic oxide (BFI) powder or use
Shake lotions (R 14 15 page 100)
b Camphor Spirit N.F.
c Benzoic Tincture U.S.P. (R 47 page 105)
- 2 One of the corticosteroids in ointment form 1-2 1/2% locally applied may be of value. The mast not be used for dendritic keratitis
- 3 Chlorotetracycline U.S.P. (Aureomycin®) 0.5% locally as eye drops may be of value in patients with dendritic keratitis
- 4 If there is associated cellulitis and lymphadenitis apply cool compresses
- 5 Treat stomatitis as outlined on page 261
- 6 Use x-ray therapy in selected cases. This requires administration by expert personnel

HERPES ZOSTER (Shingles) (code No 13 167)

An acute vesicular dermatitis of viral origin which has a characteristic distribution corresponding to involved peripheral nerves and is associated with various local nervous symptoms (neuralgia pruritus burning and autonomic sensory motor disturbances). The intercostal nerves of the sensory root of the extremities and the ophthalmic nerves are most commonly (individually and unilaterally) involved but the distribution may be generalized (resembling chicken pox). The condition is usually self limited and nonrecurrent although at times a persistent neuralgia may

ema n The d ea n i y b p r e c p t t d b y r m y b e a m n i f e s
t i o n o f c h f t i o n s l o c a l t r a u m a h e a v y m e t l p o i s o n i n g
o l y m p h o b l t o r m s

T e m tA G n l M e s

- 1 S d a t B b i t r a t e o r h o m i d e s m a y h e l p c o n t o l t
i o n d e r v o u s e s s o c i e t e d w t h n r l g
- 2 A a l g i c A e t y l s l i c y l i c A d U S P (a p i r) 0 6 5
G m (1 0 g r) o r a s p i d m p u n d w t h o r w i t h o t C o d i n e
P h s p h t e U S P 3 0 m g (1 / 2 g r) u s l i y o t o l p a r n
- 3 A t h e m o t h r a p y 1 0 c o f t h p a t e i s v o u s b l o o d i s
j e c t d i t a g l i e l i y e v y o t h e r d a y f o 3 i j e t i o n s
- 4 O p h t h a l m l o g i a l o n l i t a t i o n s h o l d b c o n s d e d f s u
p a o b i t a l i v o l e m e t t o a o l d s r o o u l s c o m p l i a t i o

B L o c i M s s

- 1 W t d s s g m y b e c s a y f o a u t a n d e x t s i v
i f l m m a t o r y l s s (e e p g e 9 8 9 9)
- 2 C a l m i n l o t o r t h e h k e l o t i o n s (s e p a g 1 0 0) a
o f t e n f a l e A p p l y l o t n l i b r a l l y a d c o v e r w i t h a p r o
t e t l a y e o f c t i o b t l i g A b o i d g r e a s
- 3 X r a y t h a p y g v n b y x p r t m y b h l p f u l
- 4 R e p o i t r y C o t c o t p i n l j e t i o U S P (c o t i c o t r o p u n
g l) 4 0 8 0 u t l M d a i l y f o 3 d y s m y l v t h e p m

LUPUS ERYTHEMATOSUS (cod No 11019)

D g

A a t r h o n i c d m t t o f k n w o i g n m n f a t d b y
t w o m n c i n i l t y p s

A D i o d T y p M l d l c a l h o p t o o s d
h k { b t i f l y p t t e r } w i t h o o s t t t a l s y m p t m s

B D i s m i t e d T y p A s e o u s s y s t e m d i s a w h a c o r m
t a d h o n f m w t h w t h o t d s o d s k j j e
n s a d a s o c a t e d w t h f v e r w e k a m i a n d v
d e n o f d i f f s l a r l s o s c h s e d c a r d i t i t h
t a n d p h r u (S p g e 5 1 9 f o r d g n i d t t m n t
f i b d e m t d t y p e)

T a t m tA G a l M s s

- 1 P f o m m p l e t m e d c a l a t d y t o r u l e o i y e t m i c i p u
y t h m t o s s
a E a m i f o c h o n i n f e c t n
b D e t m i n d e i a n d j o t t a t
- 2 P v i d p t e t i n f o m s l i g h t a n d a l l o t h e p o w f u l
d i t n D o n t u s e a n y f r m f r a d i a t i o n t h e r a p y
- 3 M i t i i o p t m a l g a l h e a l t h b y w l l b a l a n c d d i w t h
p p l e m t y i m i d r o n a d i t d f i u r d
q t t d p r c r b b d a t w h t h p a t t i f b l i
- 4 N o e p e f t h p y f o r d i s c o i d t y p e o n l y
Q u i H y d o h l o d U S P (A t a b i n ®) 0 3 G m
(5 g r) a l l y d i l y f 2 w k s t h 0 1 G m (1 1 / 2 g)
d i l y f 3 m o t h o m
b C h l o q P h o p h t U S P 0 5 G m (7 1 / 2 g r) d i l y

64 Skin Infections

for 1 week then 0.25 Gm ($3\frac{3}{4}$ gr) daily watch for signs of toxicity with both of these drugs

- c Hydroxychloroquine sulfate (Plaquenil®) 0.2 Gm (3 gr) b i d orally may occasionally be effective when quinine and chloroquine are not tolerated

- B Local Measures Treat the existing stage of dermatitis by appropriate measures (see pages 83 and 96-97)

INFECTIONS OF THE SKIN

ACUTE SUPERFICIAL INFECTIONS

The acute superficial infections include the following

- 1 Impetigo contagiosa (code No. 111.10)
- 2 Ecthyma (code No. 110.105.1)
- 3 Sycosis barbae (code No. 161.105)
- 4 Acute infectious eczematoid dermatitis (code No. 110.100.5)
- 5 Simple superficial pyoderma (code No. 1.100.1)
- 6 Secondary infections of other dermatoses

The offending organisms are usually hemolytic *Staphylococcus aureus* and/or the streptococci

Treatment

- A General Measures Systemic anti-infectives may be tried if the skin infection is resistant to local treatment if it is extensive or severe and accompanied by a febrile reaction if it is complicated or if it involves the face and danger areas (e.g. area of upper lip, nose and eyes) (See pages 486-514)

Penicillin in daily doses of 300,000 units I.M.

(CAUTION) is convenient and effective for this purpose but may be modified in dosage. Other antibiotic drugs may be substituted as the individual case demands (see page 514)

B Local Measures

- 1 Cleanse area gently with mild solution of soap and water
- 2 Soaks or compresses to involved area 15 minutes b i d (see pages 88-99)
- 3 When skin is softened by soaks gently open larger pustules and trim away necrotic tissue
- 4 Local anti-infective agents are of proved value. These may be tried individually until effective agent is determined allowing 3-4 days for evaluation. They should be applied initially at night and protected by dressings. Soaks should be applied during the day. After the area has healed any of these preparations may be applied 2-4 times daily.
 - a Neomycin Sulfate U.S.P. 0.1% in water locally q d
 - b Iodochlorhydroxyquin U.S.P. (Vioform®) 3% locally b i d in cream or ointment form
 - c Other antibiotics alone or in combination as ointments locally b i d to q i d. These include oxytetracycline, chlortetracycline and polymyxin B; combination with bacitracin or oxytetracycline, neomycin, chloramphenicol and erythromycin (see page 514)

5 Lo al age : a e of al e In c ta n c s s but attended by a
res ed risk of en tization r ct ns P icill and self
thi role sho ld not b used in outm nt form

Phylogeny

Correct pronunciation of systemic eruptions
(g d bet) o local eruptions (g m harical or hemical skin
if tations d charges et)

CHRONIC SECONDARY INFECTIONS

Date min il possibl fa tors favo ing chroni ty Obt La
bacterial cultur s and def rmin o gani m sensitivity to antibioti
agent when ver pos ibl

T h t

A Gen 1 M or

2. Consider use of vigorous systemic anti-infective therapy

B Local M e c

- 1 Us lo lmes ure as fo ac te s perfect l infection
2 T t underlying d matosis ac ording to tage and type f
le lo (s pagea 83 and 88 87)
3 Consid x aythe py if ll othe m sures ar in ffective
This must be es v d fo the specialist

ACUTE and CHRONIC INFECTIONS of SKIN APPENDAGES

Examine for local and yet milder cases of these infectious particularly if they be massive or chronic. The following disorders are included:

- 1 Folliculiti pustula (code N 181 yx2)
2 F runculosi (od No 181 100 0)
3 Carbuta le (cod No 18 100 3)
4 H dr dentitis (code No 152 100)

Limit

A Gen r 1 M ed Use vigorous systemic anti infective therapy

apply if l o s eve e t n s e mply d d o located
ind g a es (ab t e k d h d) Tt yeline USP
250 mg by mouth d ily fo s c l w e k o a t iples lfo
amid (T f yl[®] Tri orb ul[®]) t blet d ly mayb
t ed f r h n i f t n

B Loc 1 M y e

- 1 Avoid o manipulation f inflam d a e
2 U moi to dry he t to help l g le ins loc liz
3 U e p op r rgical inci l p lition or d b id m t
after l slons a e mature

COMPLICATIONS OF SKIN INFECTIONS

If pathogenic bacteria from infections of the skin invade deeper structures one of the following may be produced and other more serious infections may also occur

- 1 Cellulitis (code No 18 100)
- 2 Acute lymphangitis (code No 54 100 1)
- 3 Acute lymphadenitis (code No 55 100 1)

Treatment

A General Measures

- 1 Bed rest with immobilization of affected extremity or part
- 2 Systemic anti-infective agents must be administered in effective doses (see page 514)
- 3 Analgesics as necessary for pain (see page 32)

B Local Measures

- 1 Immobilization of affected part in slightly elevated position
- 2 Local heat to area using warm moist compresses if abscesses or pustules are present. Avoid maceration of skin (use no occlusive covering)
- 3 Local anti-infective agents to open infected areas at night

FUNGAL INFECTIONS OF THE SKIN

GENERAL CONSIDERATIONS

Diagnosis

Usually based on

A Characteristics and Location of Lesions (See below)

B Laboratory Examination

- 1 Direct demonstration of fungi in 10% potassium or sodium hydroxide preparations of scrapings from suspected lesions
- 2 Cultures of organisms
- 3 Skin tests are not reliable except that a negative reaction to Trichophyton has exclusive value when a dermatophytid is under consideration
- 4 Staining of histologic sections with periodic acid-Schiff technique

Treatment

A Local Measures

- 1 Treat acute cutaneous fungal infections initially as if acute dermatitis (see page 93). It may be necessary to treat the dermatitis before instituting fungicidal treatment
- 2 Most fungicidal agents are extremely irritating. AVOID over-treatment

B General Measures and Prophylaxis

- 1 Keep skin dry. Moist skin favors fungal growth
- 2 Cool climate where excessive perspiration and activities in hot weather
- 3 Dry after bathing and change of other clothing

e Sandals open toed shoes should be worn as they permit adequate drying of feet

f Scurf of skin should be reduced or controlled

(1) General systemic measures

(a) Sedatives to ease patients Phenobarbital
U S P 15-30 mg (1/4-1/2 gr) tid to qid

(b) Anhidrotic drugs (e.g. atropine) are usually ineffective

(2) Local measures

(a) Talc or other drying powders (see page 99)

(b) Drying oaks (see pages 98-99)

g Toughen skin by gradual daily sunbaths or by quartz lamp treatment

2 Foci of fungal infections should be eradicated or controlled
a Trunk, navel, umbilicus, groin, webs of toes and other areas where fungi are found

b Group of community showers or bathing places unless strictly supervised should be avoided

TINEA CAPITIS (Ringworm of Scalp) (code No. 162.211)

This contagious dermatomycotic condition occurs almost exclusively in children. It is very persistent but less spontaneous than the scalp type. The lesions are originally red and scaling and eventually form a mass of alopecia. Fluorescence under Wood light is characteristic in Microsporum infection (90% of cases in some areas). There is often a history of contact with infected individuals. Household pets

Treatment

A General Measures No

B Local Specific Measures It may require 2 months or more to cure the disease. The human type is more difficult to treat than the animal type (dogs and cats).

1 Scalp cleaning and preparation (Netschell)

a Clip hair closely every 2 weeks and have patient wear clean stocking cap or klick cap for protection

b Wash scalp as necessary

2 Fungicidal astringents Rub the well into scalp morning and night after scalp has been washed

Selenium sulfide NF 5% in Cbow 1500[®] ointment (polyethyl glycols)

b Benzoin acid and salicylic acid ointment (Whitefield's one-half strength) (R 34 page 104)

Sulfur 12% in ointment (R 35 page 104)

3 Epilation Use Waxing or 250 Wt purple X-1 mop to remove infected hairs daily by tweezers by adhesive tape treatment

4 X-ray may be used effectively and may work when hair is removed mechanically as a fall X-ray therapy may be given by trained personnel only. Do not re-epilate with x-rays

Prephylaxis

A Individual

1 Exchange of hairgear must be avoided

88 Versicolor and Corporis

- 2 Infected individuals or household pets must be vigorously treated and re-examined for determination of cure
- 3 Scalp must be washed after barber shop haircut

B Group

1 Routine school surveys may be advisable

2 Epidemic precautions

- a Wood light examination of students less than 12 years old
- b Isolation of infected individuals in special classrooms
- c Careful follow up of infected individuals and periodic re-examination of all children until all cases are cured
- d Education of barbers regarding handling of infected individuals

PITYRIASIS VERSICOLOR OR TINEA VERSICOLOR

(code No 11 208)

A mild condition characterized by tan or pinkish erythematous macules of variable sizes mildly pruritic usually on the upper trunk. Healed areas remain depigmented for a few months. Coarse brown hyphae and large spores in clusters may be demonstrated easily in skin scales prepared with 10-15% sodium hydroxide.

Treatment

A General Measures Encourage no mal skin hygiene

B Specific Measures One of the following may be used

- 1 Sodium thiosulfate 10% aqueous solution b i d
- 2 Mild Whitfield ointment 1/4 1/2 strength (S 34 pag 104) at bedtime

TINEA CORPORIS OR TINEA CIRCINATA

(Body Ringworm) (code No 130 211)

Body ringworm is characterized by single or multiple (relative ly few) scaly papules circular in shape with clear central areas and with minute vesicles in the actively spreading periphery they are found most commonly on the trunk neck and limbs Lesions occur occasionally as thick pigmented patches Diagnosis should be confirmed by demonstration of the fungi

Treatment

A General Measures (See page 86)

B Local Measures Avoid overtreatment

1 Treatment proper stage of the dermatosis (see 83 88 97)

2 Fungicidal agents

- | | | |
|----------------------|------|-------|
| a R Salicylic acid | 0 3 | gr v |
| Sulfur ppt | 0 8 | gr xv |
| Hydrophilic ointment | | |
| q x ad | 30 0 | 3j |

S g Locally b i d

- b Compound Undecylenic Acid Ointment N F may be used in the less chronic and nonthickened lesions

Prophylaxis

1 General Measures on page 86

- a Avoid contact with infected household pets
- b Avoid removal of clothing without adequate laundry

TINEA CRURIS (Inguinal Ringworm or Jock Itch) (code No 146 215)

Erythematous macular lesions with sharp margin, cleared center and thin spreading peripheral rim in intertriginous areas (chafing or friction areas) such as groin, scrotum and axilla. The fungus should be demonstrated in differential condition from athlete's dermatitis.

Treatment

- A General Measures** See general rules (page 86) but also
- 1 Drying powder (page 99) should be dusted into involved areas 2-3 times a day especially when perspiration increases.
 - 2 Keep the area clean and dry but avoid overbathing.
 - 3 Prevent intertrigo or chafing by avoiding over-treatment of the perianal area to further infection and complicate treatment.
 - 4 Clothing Avoid rough textured clothing.
- B Local Measures**
- 1 Treat the edges of dermatosis (see page 93). Secondarily infected or inflamed lesions are best treated with petrolatum ointment applying soothing and drying solution to involved areas. Use wet compresses of Potassium Permanganate U.S.P. 1:10,000 (or 1:20 Aluminum Acetate Solution U.S.P.) or in case of anogenital infection to lesions by tub baths.
 - 2 Fungal dermatitis
 - a Self-examination (see page 101)
 - b Wet solution of iodine (not more than 1% tincture) boric acid Calomel Fuch's Solution N.F. (Castile paint) 1/3 strength on a day.
 - d Compound Sulfur Acid Chlorine N.F. boric acid Sulfur lye and intimate (see page 104)

DERMATOPHYTOSIS (Tinea of Palms and Soles) (code No 112 211)

A relatively common athlete's foot dermatosis occurring on the sole of palms and interdigital areas with a characteristic cyrtotic curvature. Acute reddened weeping vesicular lesions are seen in the acute stage. (Vesicular lesions of the feet are most commonly due to fungi but on the hands are more commonly due to other causes.) Contact dermatitis but also infections and reactions. The area is slightly fissured and macerated lesions in the subungual tags and beneath and around the nail in the broad tips. Diagnosis should be confirmed by demonstration of fungi.

Treatment

- A General Measures** See General Measures (page 86) but put particular emphasis on personal hygiene
- 1 Rubber or wooden sandals should be used in community showers and bathing places.
 - 2 Open toed shoes and sandals are better for general wear.
 - 3 1% Sodium Hypochlorite Solution N.F. foot soaks before and after bathing in community showers or of bathtub is recommended. Dry between toes after shower.

88 Versicolor and Corporia

- 2 Infected individuals or household pets must be vigorously treated and re-examined for determination of cure
- 3 Scalp must be washed after barber shop haircuts

B Group

- 1 Routine school surveys may be advisable
- 2 Epidemic precautions
 - a Wood light examination of students less than 15 years old
 - b Isolation of infected individuals in special classrooms
 - c Careful follow up of infected individuals and periodic re-examination of all children until all cases are cured
 - d Education of barber regarding handling of infected individuals

PITYRIASIS VERSICOLOR OR TINEA VERSICOLOR (cod No 112 208)

A mild condition characterized by tan or pinkish very fine scaly macules of variable sizes mildly pruritic usually on the upper trunk. Haled as a rem in dep gm t d for a f w mo the Co rse blunt hyphae and large pores a clu t s may b dem n str ted easily in skin s l s prepared with 10-15% of m hydro ide

Treatment

- A Cerl Measure Encourage normal skin hygiene
- B Specific Measure One of the following may be used
 - 1 S d un thio sulfate 10% aqueous solution b i d
 - 2 Mild Whitfield's ointment 1/4 1/2 at gth (R 34 p 96 104) at bedtime

TINEA CORPORIS OR TINEA CIRCINATA (Body Ringworm) (code No 130 211)

Body ringworm is characterized by single or multiple (relatively few) scaly papules circular in shape with clear central areas and with minute vesicles in the actively spreading periphery they are found most commonly on the trunk neck and limbs. Lesions occur occasionally as thick pigmented patches. Diagnosis should be confirmed by demonstration of the fungi.

Treatment

- A General Measure (See page 86)
- B Local Measures Avoid overtreatment
 - 1 Treat the proper stage of the dermatosis (see 93 96 97)
 - 2 Fungicidal agents
 - a Salicylic acid 0.3 g v
 - Sulfur ppt 0.9 gr xv
 - Hydrophilic ointment
 - qs ad 30.0 31
 - Sig Locally b d
 - b Compound Undecyl nic Acid Ointment N F may be used in the last 4 months of the disease

Prophylaxis

- A General Measures on page 86
- B Avoid contact with infected household pets
- C Avoid exchange of clothing without adequate laundering

TreatmentA G LM sa r NoneB L al M a r a

- 1 M n cal Sa dpaper o fil daily (down to nail bed if necessary) Srg cal avulsio of th n ll m y be necessary
- 2 F glc d i gents Apply on i fect d n tis
 - a Iodine Tincture U S P 0 1 1 0% b i d
 - b Chrysarobin 4% i hlo form b i d
 - c Ch y robin U S P 0 1 0 5% in Petrol tum U S P b i d
 - d Whitfield s o tme t 1/2 strength b i d (§ 34 p ge 104) Diamth ole D hydro hi r d N N D (Aste ol®) o i t m t 5% locally b i d
 - f Ve defam® liquid (o d i m prop ionat s d um aprrylate prop i acid und yleni cid s licylic acid copp u decyl nat) applied b i d
- 3 X y f ctio al dos (gl o ly by t ain d person el) may be of aid in m ld cases and m y requ e m nths for c re Some a thorit feel th t x y ha o plac in the tre tme t of o ychorny is

P phyl

See Gen l Co derat na p g 86

INFESTATIONS OF THE SKIN

SCABIES (ode No 110 281)

A mmo derm tit s ed by i festation with Sa opt s a b nd h t i d by m ll p u t s i s pust les s d c i t on fo d most f equently in the f g r w ba body flex butt cks nippl nd g nit lia Th f m l mit can be d m tated th ki leal Scabies is f u d m o t i f que tly u d nhyy c d t ons with h t ry of po r to ab s Thj i f t ti es ma to b dia pp ng in th U S A

T tm t

- A G LM sa sa N cid i d unl as seve es cond y prod m i p ent This m y eq ur yat mi ti f cti g nt pr i o with th ants bel tr atm t
- B Loc LM a r U le s i ons ar compl ated by e re se o da y py d m t eatm ti d ect dp m ly t wa d di i f st ion If s o d ry pyoderm p t P lass m Pe mang te U S P so ka (1 10 000) 1/2 h b i d i t i d m y b indicated b fo defi t v i tm nt
- C D f tlv M a
 - 1 G mm Be e He a hlo id U S P (Gamm ® Kw 11®) 1% nce mbs 2 o appli d a h nght for 3 ght (§ 40 p g 104) It m y b d th p e e o f i f t on It is n w id d to b th t alme t f ch ic
 - 2 S lfu t alme t (do l m thod now ob o i t) P l m i a y b thng H p ti t ak a h w nd cr b vigo o ly with hot s p d w t b S H o tm t 4 5% r bbed in th o ghly f om k d wn to t e h nght f 3 5 s c v nght g l d b thng f i t t p i t ot to ch ge

- 5 Socks should be changed frequently
- 6 Dusting and drying powders p r n (see page 89)
- 7 Place small wads of cotton between toes at night

B Local Measures Do not overtreat

- 1 Acute stage (This will vary from 1 to 10 days) Treat as for any acute dermatosis using soaks (see page 88) for 20 minutes b i d or t i d If secondary infection is present use 1:10,000 Potassium Permanganate U S P soaks If secondary infection is severe or complicated treat as per directions on pages 84-86
- 2 Subacute stage Any of the following may be used
 - a Zincundecate ointment b i d
 - b Whitfield's ointment $\frac{1}{4}$ to $\frac{1}{2}$ strength (R 34 page 104)
 - c Sol. of coal tar 5% in starch lotion or R 17 page 100
 - d Coal tar 1-2% in Lassar's paste
- 3 Chronic stage Use any of the following
 - a Iodine as 0.1-1% tincture paint on areas daily
 - b Whitfield's ointment $\frac{1}{4}$ to $\frac{1}{2}$ strength (R 34 page 104)
 - c Compound Uricolytic Acid Ointment N F b i d
 - d Alcoholic Whitfield's solution (R 46 page 105)
 - e Carbol Fuchsin Solution N F (Castellani's paint)

C Manual Measures Remove or debride dead or thickened tissues after soaks or baths by careful manual technique

D X-ray therapy may be of value when hygienic and chemical means fail Must be given only by trained personnel

DERMATOPHYTIDS (Allergy or Sensitivity to Fungi)
(code No. 111 2115)

An erythematous vesicular or eczematoid dermatitis of hands (and less commonly of other skin areas) occurring secondarily to a recent fungal infection and pathologically following over-treatment of such infections. The skin reaction will vary in severity with the activity of the local organisms. Fungal elements present in the primary lesion but are not present in the secondary lesions. Trichophyton 0.1 cc intracutaneously holds give a positive reaction (it is read in 24 to 48 hours like a tuberculin test).

Treatment

A General Measures None

B Local Measures

- 1 Treat lesions according to type of dermatitis (see pp. 96-97)
- 2 Treat primary focus as indicated

Prophylaxis

- 1 General Measures on page 86

TINEA UNGUIUM OR ONYCHOMYCOSIS
(code No. 170 2)

A destructive condition of one or more (but rarely all) fingernails or toenails which begins at the lateral borders and often eventually causes deformity and even separation of the nail plate. Diagnosis should be confirmed by demonstration.

GENERAL RULES GOVERNING CHOICE OF TREATMENT OF VARIOUS STAGES OF DERMATOSES

Recomm d M d ments		Page
<p>ACUTE LESIONS Charact tics R t onset r d burning swollen it hng bll te lng and o f g</p>	Soak Fo lesions l l d to xtr mitties	98
	Wet d aings F localiz d l sio s of h d neck t u k o e t r e m i t t i s	98
	B ths F o r g e n e r l l e d l e s i o n s	68
	Shak lotions	100
<p>SUBACUTE LESIONS Ch acteristics Int m d i t e d u r a t i o b l d i n g l i and l g r y i n p p a n c e</p>	Em l i o	100
	Hyd ophil O t m t	103
	P a t (high powd r o tent)	103
	Cold ams	103
	Cr ams (ontain water)	103
	G e a s y O u n t m t	104
<p>CHRONIC LESIONS Char t lctics L o g e r d r t i n q t t h l k n e d e n r u t e d f i s r e d and scaly</p>		

Exact directions for choice of treatment will vary with the individual case. This will depend upon a wide variety of factors including characteristics of the dermatosis, extent of lesions, general character of patient's skin, previous medication and drug allergies.

- clothing or bed linen and not to bathe during this period
- d Final bathing On the day after the final successful rub-in instruct the patient to take another scrubbing with hot soap and water in a shower and to change to all clean personal clothing. Bed linen must also be changed
 - e Soothing lotions It is often necessary to prescribe soothing baths (see page 86) and shake lotions (§ 14, 15, 16, page 100) frequently following the treatment in certain patients with sensitive skins
 - f Personal clothing and bed linen must be laundered or cleaned

Prophylaxis

- A Good hygiene
- B Avoid intimate contact with infested individuals

PEDICULOSIS

Name of Disease

Synonyms

- 1 Pediculosis pubis (code No. 110.292) Pubic louse (c. abs.)
- 2 Pediculosis corporis (code No. 110.2912) Body louse
- 3 Pediculosis capitis (code No. 110.91) Head louse

Diagnosis

Diagnosis is dependent upon demonstration of lice or nits (eggs) with evidence of pruritic dermatitis. Scratch marks and pyoderma often tend to obscure primary puncta.

Treatment

- A Definitive Treatment 10% Chlorophenothan U.S.P. (DDT) in talcum or pyrophyllite is extremely effective in all forms of pediculosis
 - 1 Scalp Dust 2.5-5.0 Gm (37 1/2-75 g.) well into scalp and distribute evenly over scalp. In treatment patient not to wash hair for 1 week. Repeat treatment at end of 2 weeks
 - 2 Body Either powder or spray can be used
 - a Dust powder evenly over body surface. May take 5-10 Gm (75-150 gr.)
 - b

Chlorophenothane (DDT)	6.0 g.
Ethyl aminobenzoate (Benzocaine®)	12.0 g.
Tween 80®	14.0 g.
Benzyl benzoate q.s. ad	100.0 g.

 Sig: Dilute 1 part of this solution with 5 parts of water. Spray all hairy parts of body with about 20 cc of liquid. Protect the eyes. (CAUTION: Benzocaine® is a sensitizer.)
 - 3 Pubis Dust powder liberally and distribute evenly. Allow powder to remain for 2-3 days. Wash off with soap and water. Re-examine in 1 week and reapply if necessary
- B General Measures
 - 1 Thorough bathing with hot soap and water
 - 2 Disinfection of discarded clothing in case of body louse. Autoclaving or suitable laundering methods must be used
 - 3 Hairy areas in case of pediculosis may need to be clipped
 - 4 Treat dermatitis (see pages 83 and 86-87)

Name	Action	Preparation	Technique
E Hydrophilic Ointment U S P	Vaginal cream For psoriasis or seborrhea Fungal For seborrhea	One may add 5% ammoniated mercury 1% salicylic acid 3% sulfur 5% dithranol solution of coal tar	Apply sparingly with fingering tip b i d
F 3% Slicylenilid N F in Ca bowax 1500®	Follicular capitis	Disperse 60 Gm of Slicylenilid (ointment)	Locally b i d to the scalp (It is not necessary to clip the hair)
G Gamma Benzene He chloride U S P	For scabies and pediculosis	Dispense 30 Gm or 3i	Locally b i d for 1 to 3 days
H Aqueous emulsion (0.1%)	For pyoderma	8 N omycin 0.12 gr ii Dithranol water q ad 120.0 3iv	Apply with cotton b i d to q i d
I Methyrosaniline Chloride U S P (Giant 1 let)	For monilia	Dispense 30 c or 3i of 1% aqueous solution	Paint on with pipette for o c d i l y
J Paraaminobenzoic Acid N F in methyl alcohol (10%)	Protect from actinic rays	Dispense 60 Gm or 3i	Apply to exposed surfaces achmor ing
K Lassar's Paste (Zinc Oxide Paste U S P)	Protective and soothing	Dispense 30 Gm or 3i	Locally b i d

USEFUL MEDICATIONS FOR SKIN DISEASES

Name	Action	Prescription	Technique
A Sulfur and soda baths	Cleaning and soothing	1/2 cup each of coarsely powdered soda to a tepid bath	No soap The patient is bathed for 15 minutes then dabbed (not rubbed) dry
B Cool wet dressings	Cooling soothing antipruritic	1 Tbsp to 1 qt or 1 liter of cold water	Wring out a washcloth or Turkish towel and lay on the affected areas 15 minutes twice a day or continuously Use no waterproof covering as one of the chief uses of these agents depends on the cooling effect of evaporation (Note Boric acid and potassium permanganate are poisons and should not be used internally or on large denuded areas)
b Alumumacetate USP	Cleansing mildly astringent	Domeboro® powder 1 tsp to 1 qt or 1 liter of cold water	
c Potassium Permanganate USP BP	Deodorizing	On 0.3 Gm (5 gr) tablet dissolved in 1 qt or 1 liter warm water	
C Hot wet dressings Magnesium Sulfate USP BP	Promotes blood flow Localizes infections and deep pharyngitis	2 Tbsp to 1 qt or 1 liter of hot water	Apply as above using washcloth or Turkish towel
D Sterch U S P B P (Add 5% detergent solution of castor oil)	Soothing and drying (Acetoplastin and healing)	R Sterch 36 3 x Zinc oxide 36 3 x Glycerin 18 3iv Lmewt qs ad 180 3 i	Apply with cotton ball

Type of Skin Lesion	Example	Methods of Local Treatment Always start at the stage with a type of dressing
5 Ulcer Simple superficial Deep pyogenic Dephlegmated Ulcers Simple wheals Angiodermatosis	Simple ointment Tropical ulcer Trophoblastic Hives Angiodermatosis Simple fissures	Washing, antiseptic, lotions and ointments Antipruritic soothing bath and shock lotion Sleeve, strapping, antiseptic applications and ointments (old wet)
6 Erythematous lesions Scarcely crusted Infected crusts	Eczema Impetigo	Wet dressing, debridement, follow with hot lotion and grease Wet dressing, debridement, follow with antiseptic solutions and ointment
9 Dermatitis Adhesive Non-adhesive (foliaceous) Gyrate lesions	Pustules Exfoliative dermatitis Scabies Itching	Keratolytic and later keratoplastic agents Wet dressings, baths, shake lotions, emulsions and later grease Keratoplastic agents Wet dressing, shock lotion and powder
10 Maculated lesions	Itching	

METHODS OF LOCAL TREATMENT OF VARIOUS TYPES OF SKIN LESIONS

Type of Skin Lesion	Example	Methods of Local Treatment Always treat the stage as well as type of dermatitis
1. Macule Simple erythema (asymptomatic) Burning erythema	Drug erythema Sunburn	Soothing wet dressings or shake lotions
2. Papule Maculopapular lesions Papulosquamous lesions acute chronic Anthrax lesions Lichenified lesions Verrucous lesions	Psoriasis Acne vulgaris Lichen planus Verru vulgaris	Mild keratoplastic lotions and ointments Soothing wet dressings or shake lotions Keratoplastic and later keratolytic agents Keratolytic and astringent agents Keratoplastic and later keratolytic agents Keratolytic and caustic agents
3. Vesicle Multiple vesicles or diffuse weeping lesions Herpetic lesions Bullae	Eczema Herpes zoster Pemphigus	Soothing wet dressings during daytime and shake lotions or pastes at night. As process subsides change to pastes and creams Shake lotions or wet dressings Wet dressings of blisters and shake lotions
4. Pustule Impetigo Ecthyma Furunculoid Follicular	Impetigo Ecthyma Furunculoid Syphilis barba	Wet dressings debrided anti-infective powders solutions and ointments Wet dressings of blisters anti-infective powders solutions and ointments Wet dressings of blisters (only when ripe) and drainage (caution on lesions above lip) Wet dressings anti-infective solutions and ointments

Agit	Act n	Range (C n c n ratio s U d	Mo t Common Str gth U d	Pep tion of S ul n f M t Comm ly Employ d St e gth
R 7 Mercury Bichloride N F em il po t bl t	Ant Ant ptic		1 10 000 (0 01%)	O t bl et HgCl ₂ to 1 qt o 1 l t w t Poison Do not use on denuded areas
R 8 Pot i m Pe manganat U S P R P	Antiprur t Oxidizing Ant pt A stringe t	1 10 000 to 1 400 (0 01% to 0 25%)	1 10 000 0 01%	One o 3 Gm (5 gr) t bl et KM O ₄ to 3 qt o 3 liters water or 0 1 Gm (1 1/2 gr) KMnO ₄ to 1 qt or 1 lit r water
POWDERS				
Name	P	Ap on	In tructions and Rem ks	
R 9 Absorbent Gelat Sp ge U S P (o st il l)	G I am [®] powder	10 0 Gm	Fo leg ulcers and the indole t ul ers It is absorbable hemostatic gelati Apply b i d	
R 10 Talc U S P (t leu r)			Simple du ting p wder	
R 11 Foot powder imple	R S licydic cid Bor ca d Talc m	1 0 g x 1 0 gr x 100 0 3xxv	Simple powder	
R 12 D t ing powder antipru itic	R Camphor powd r d Zinc o lde powder d Starch powdered q s ad	2 0 6 0 3 s les 16 0 3lv 100 0 3xxv	S mple antipruritic powd r	
R 13 Chlorophenothale U S P (DDT)	R DDT Talcum q s ad	10 0 3l s 100 0 3xx	Sig Apply 1/2 to 1 o over the entire ur face of underw ar and t eat s arms on inside of shirt s d trousers Remarks Effective against all p dical is	

SIMPLE SOLUTIONS FOR SOAKS AND WET DRESSINGS

Indications. For acute red swollen itching infected weeping or vesicular lesions. Solutions must be applied cool (not for infections).

- (1) Basin soaks (2 5 quarts of solution) for hands and feet 1/4 hour b i d
 - (2) Wet dressings for localized lesions use turkish towel keep saturated with solution
 - (3) Open dressings for very acute lesions and when marked cleansing and soothing action is desired
- Frequent applications are necessary (1 c for 1/2 hour b i d to q i d)
- (b) Covered dressings should not be used

All of the solutions have a drying, soothing and cleansing action in addition to those mentioned

Agent	Actions	Range of Concentrations Used	Most Common Strength Used	Preparation of Solution of Most Commonly Employed Strength
Plain tap water	(See above)			
R 1 Sodium Chloride U S P B P	(See above)	6 1 000 to 1 000 (0 5% to 1 5%)	0 9%	2 1/4 dr (2 tsp) NaCl to 1 qt water or 9 Gm NaCl to 1 liter water
R 2 Sodium Bicarbonate U S P B P	Antipruritic	1 50 to 2 (2% to 5%)	3 0%	7 1/2 dr (8 tsp) NaHCO ₃ to 1 qt water or 30 Gm NaHCO ₃ to 1 liter water
R 3 Boric Acid U S P B P	Antipruritic	1 50 to 2 (2% to 4%)	3 0%	7 1/2 dr (8 tsp) H ₃ BO ₃ to 1 qt water or 30 Gm H ₃ BO ₃ to 1 liter water
R 4 Magnesium sulfate U S P B P	Antipruritic	1 50 to 2 (2% to 4%)	3 0%	7 1/2 dr (8 tsp) NaHCO ₃ to 1 qt water or 30 Gm NaHCO ₃ to 1 liter water
R 5 Aluminum acetate Sol U S P	Astringent	1 200 to 10 (0 5% to 10 0%)	5 0%	4 Domeboro® tablets or 50 cc Burow's sol (N F) to 1 qt or 1 liter water
R 6 Silver Nitrate U S P B P	Astringent Antiseptic	1 10 000 to 1 200 (0 01% to 0 5%)	1 400 to 0 25%	10 cc of 25% AgNO ₃ solution or 2 5 Gm AgNO ₃ to 1 qt or 1 liter water

Ag t	Action	Range of Concentration	Most Commonly Employed Strength	Preparation of Solution
R 7 Mercury Bichloride N F small poison tablets	Antiseptic Antigraft		1:10,000 (0.01%)	One tablet HgCl ₂ to 1 qt or 1 liter water. Poison. Do not use on dermated areas.
R 8 Potassium Permanganate N F	Antiseptic Oxidizing Antigraft	1:10,000 to 1:400 (0.01% to 0.25%)	1:10,000 0.01%	One 0.3 Gm (5 gr) tablet KMnO ₄ to 3 qt or 3 liter water or 0.1 Gm (1½ gr) KMnO ₄ to 1 qt 1 liter water.
POWDERS				
Name	Description	Preparation	Instructions and Remarks	
R 9 Absorbable Gelatin Sponges (sterile)	Gelform powder	100 Gm	For leg ulcers and other indolent ulcers. It is absorbable hemostatic gelatin. Apply by dusting.	
R 10 Talcum Powder (talcum)			Simple dusting powder	
R 11 Foot powder simple	Sulfuric acid Boric acid Talcum	10 gr 10 gr xv 100 0 5 xv	Simple powder	
R 12 Duhring powder antipruritic	Camphor powdered Zinc oxide powder red Starch powder	20 6 0 3as 1as 18 0 3iv 100 0 5 xv	Simple antipruritic powder	
R 13 Chlorophenothene N F (DDT)	DDT Talcum	10 0 3ias 100 0 5xv	Sig. Apply 1/2 to 1 z over the entire surface of undrained and treat areas on inside of shirt and trousers. Remarks: Effective against all pediculosis.	

LOTIONS AND EMULSIONS

Liquid mixtures containing ingredients in solution and/or suspension Useful in a wide variety of localized and generalized skin lesions because of ease of application and removal They have a marked drying effect and must be avoided if this effect is undesirable The following are some useful well known lotions

Lotion and Action	Prescription	Instructions and Remarks
R 14 Calamine Lotion (L S P (soothing drying))	R Prepared calamine 10 0 3 iss Zinc oxide 10 0 3 iss Glyc erin 2 5 3 ss Magnesia of bentonite 51 0 3 v liss Lime water q s ad 1 5 0 3 xxxl	Sig Apply to affected area q i d or p r n Remarks Use for acute dermatitis Avoid excessive drying by prolonged use of this lotion (as with other non oily lotions) Add 1% pt nol for antipruritic effect
R 15 Starch lotion (antipruritic soothing drying)	R Starch corn 24 0 3 vi Zinc oxide 24 0 3 vi Glyc rin 1 0 3 iii Lime water q s ad 120 0 3 iv	Sig Apply locally b i d and p r n Remarks Use for acute dermatitis Useful basic lotion to which other agents may be added
R 16 Oily lotion (soothing drying lubricating)	R Zinc oxide 10 0 3 liss Olive oil Lime water ad q s ad 120 0 3 v	Sig Apply to affected area q i d or p r n Remarks Use for acute dermatitis Less drying than R 14 and 15
R 17 Coal tar lotion (soothing drying keratoplastic)	R Sol coal tar 12 0 3 i Zinc oxide 24 0 3 vi St eb 24 0 3 i Glyc rin 36 0 3 ix Water q s ad 120 0 3 iv	Sig Apply locally at night Scrub in a m Remarks Use for subacute dermatitis Useful when itching lotion
R 18 Sun screen lotion (protective)	R Para amin benzoic acid 3 0 3 i Emulsion base q ad 30 0 3 i	Sig Apply locally to skin before each exposure to the sun

Lotion and A t t n	Pre s i p t i o n	In t r u t t l e s a n d R e m a r k s
R 19 Acne lotion	R Sulf ppt 3 6 3i Zinc sulf 1 -- Sodium borate Zinc oxid 3 A to c 30 0 3i Camph r water Rose water 32 q s ad 120 0 3i	Sig Apply loc ally t night R m ke For acne
R 20 Sulfur reso lnel lotion (d ying antipruriti fungi cid l k r tolytic)	R Sulf ppt 4 0 1 Resor l 1 2 0 3 s Zinc ide 25 0 3vi Talc 25 0 3vi Bento lite 5 0 3i Al ohol 50% q s ad 120 0 3i	Sig Apply lo ally at night Se ub in a m Remarks For subacute and chronic dermatitis The sulfur and resorcinol conc tration may be do bled or t ipled if more stimulating effect is d ired
R 21 T scalp lotion (kerat pl stic)	R Sol c altar 20 0 3v Cast r oll 8 0 3ii Alcoh 1 95% q s ad 120 0 3iv	Sig Rub small quantity into scalp at night R em a k s All purpose scalp loti n
R 22 Me ury allyclic hair lotio (k atopl stic)	R Me ury bichloride 0 1 gr les Sali ylic acid 3 0 gr xiv Alcohol 50% q ad 120 0 3iv	Sig Rub small q antity into calp at night R marks All purpose scalp loti
R 23 Und rarm totl n (antip pl ant)	R Al minum chl ide 50 0 3ii Gly erin 30 0 3i Distilled water q s ad 240 0 3viii	Sig Apply sm ll quantity to u derarms ea h m rning R marks U eful antipe spirant

OINTMENT BASESIndications.

- 1 To correct fat deficiency in a dry skin
- 2 To provide mechanical protection to the unduly lesion
- 3 To help absorb or imbibe transudates from underlying lesions (This holds true only for the hydrophilic preparations)
- 4 To apply active medicinal agents to the skin

Contraindications.

- 1 Acute inflamed sores
- 2 Hairy areas (except the hydrophilic preparations)

Preparation		Preparation	Pharmacological Properties
OINTMENTS			
R 24	Petrolatum White U S P White Soft Paraffin B P		Chemically inert Retards penetration of incorporated medicaments in some cases
R 25	Petrolatum Hydrophilic U S P	3% cholesterol in white wax and steryl alcohol	Favors penetration of incorporated medicaments Imbibes water (hydrophilic)
R 26	Wool Fat Hydrocarbon U S P B P (lanolin)		Adheres well to skin stable favors penetration Water sensitive
R 27	Wool Fat U S P B P (a hydrocarbon)		Imbibe water Favors penetration Water sensitive
R 28	Zinc Oxide Ointment U S P B P	20% zinc oxide in liquid petrolatum wool fat wax and white petrolatum	Chemical protection Imbibe water stiffens ointment gives body to ointment Makes it stick
R 29	Thiodibromide U S P B P (Cocobutyl)		Melt at body temperature

P P t		P		Ph m log		P op t	
CREAMS (C t n w t e)		Ad ant g		pt		outm nt	
R 30	Hyd ophall U S P	R M thyp rab		0 025 g		3 6	
		P opyba ben		0 015 gr		1/4	
		St a yl al olol		25 0		3 i	
		White p t ol tum		25 0		3vi	
		Pr pyl egly l		12 0		3ii	
R 31	R s w t U S P	Polyoxyl 40 stearate		5 0		3 1/4	
		P f d w t r q ad		100 0		3 v	
		R Sperma cl		12 5		3i 1	
		Wht w x		12 0		3iii	
		Expr ed alu nd oil		56 0		3xi	
R 32	Em l st n b se	S dium b te		0 5		gr viles	
		Ro e wat		5 0		3ii 1/4	
		D still d wat		14 0		3iia	
		Ro oil		0 02		3i 1/3	
		R Dup 10 C		1 6		3 v	
PASTES (High p wder nte t)	Z Oxide Past U S P (L s r P te)	C tyl alcohol		7 0		3ii 3/4	
		St yl l hol		7 0		13/4	
		White p tr lat m		20 0		3v	
		H avy liquid petrol tum		2 0		3	
		Rutob n		0 05		3 3/4	
R 33	Z Oxide Past U S P (L s r P te)	Di ill d w te q s ad		100 0		3 xv	
		Promote ev potat n s d cooling d c		see v alc l t n			
		R Zin xide		25 0		3vi	
		Sta h		25 0		3vi	
		P t olatum w hte q ad		100 0		3 xv	

Mechanical prot tive Increas
adhe ion b t d c ess spe t ation
of m dicam nts (Add 2% choles
terol or 5% c tyl alcohol to increase
water limbing power)

Cold c m (wat n oil) cooling
nd soothing eff t

Non h sting and n ritat g L
messy than oth r creams and out
m nts

OINTMENTS, MISCELLANEOUS STANDARD PRESCRIPTIONS

Common Name	Prescription	Preparation	Instructions and Remarks
R 34 Ointment of Benzoic and Salicylic Acid U S P (Whitfield's)	Benzoic acid Salicylic acid Polyethylene glycol ointment q s d	60 0 100 30 0 30 100 0 300	Sig Apply locally to skin at bedtime Remarks Effective fungicide. Often best prescribed in 1/4 strength. Not for acute or subacute lesions.
R 35 Aluminate ointment (123)	Aluminate Sol N F Wool fat Zinc oxide paste	10 0 300 10 0 30 10 0 30	Sig Apply locally to skin p r n Remarks Valuable on receding inflammatory process.
R 36 Sulfur salicylic acid ointment	Sulfur Salicylic acid Petrolatum q s d	10 0 30 gr xv xlv 10 0 30 gr xv xlv 100 0 300	Sig Apply locally to skin p r n Remarks Potent fungicidal combination. 10% FOR ACUTE OR SUBACUTE LESIONS.
R 37 Calamine cream	Hydrophilic ointment U S P Calamine lotion	33 0 300 66 0 300	Sig Apply locally to skin p r n Remarks Good general purpose cream. Useful vehicle for water soluble agents.
R 38 Ammonio Mercurio Ointment U S P	Ammonio mercurio Liquor petrolatum Petrolatum q s d	50 0 gr 1 xv 30 0 30 100 0 30	Sig Apply locally to skin p r n Remarks For seborrheic dermatitis and psoriasis.
R 39 Kaolin and sulfur ointment	Kaolin Sulfur ppt Zinc oxide ointment q s d	10 0 300 10 0 300 100 0 300	Sig Apply locally at bedtime Remarks A good substitute exfoliating paste for acute.
R 40 Calamine Benzene Hexafluoride U S P (Kew)	Kewell's ointment	60 0 300	Sig Apply as directed Remarks Useful as abradant.

SOLUTIONS TINCTURES AND PAINTS

Name	Preparation	Remarks
R 41 Methyl Chloride USP (Gentian violet)	10% solution	Antipruritic (gentian violet)
R 42 Sulfur USP	10% solution	Fungicidal (pityriasis)
R 43 Silver Nitrate USP	10% solution	Ulcericidal (giant cell)
R 44 Zinc USP	4% solution	Fungicidal (pityriasis)
R 45 Nitrogen (Methyl)	0.5% (1:200) solution	Bactericidal and germicidal agent
R 46 Alcohol White Lotion	8% solution Benzoin 2.0 g Alcohol 4.0 g Aqueous 120.0 g	Apply locally to skin Effective fungicidal combination May substitute by rum for alcohol
R 47 Boric Compound Tincture	8% solution	Ulcericidal (giant cell)
R 48 Sulfur Lotion USP	USP 65% solution of 8% (oil)	Fungicidal (pityriasis)
R 49 Tincture Lotion	8% solution Oleic acid 4.0 g Methyl 8.0 g Methyl 100.0 g	Add up to full capacity of water to make shampoo
R 50 Nystatin (Mycostatin)	8% solution Oral tablets (500,000 units per tablet) Vaginal tablets (100,000 units) Oral tablets (100,000 units per Gm) Dusting powder	Tincture of whole (fungal disease)

DERMATOLOGIC MEDICAMENTS

The following drugs may be incorporated singly or in combination in the lotions and ointment bases listed on pages 100 and 102. In general, it is preferable to make preparations as simple as possible. The pharmacological action of the various drugs depends not only on the inherent chemical characteristics of the agents but also upon their concentration. (E.g., lactic acid is astringent at 2-3% but at 4-6% and is cytolytic at 6-10%.) Often, even a little drug may

be used to achieve a common desired result. Preference for certain drugs may be based on actual superiority or only upon tradition. Agents and concentrations employed will depend upon the clinical characteristics of skin lesions and upon individual variations in tolerance. It is usually desirable to begin with weaker concentrations and to increase strength as indicated.

*Cannot be prescribed in cream bases only in the hydrophilic bases (e.g., Aquaphor® Eucerin® Carbowax 1500®)

Type of Drug	Name of Drug	Concentration on Employed
ASTRINGENT and/or DRYING (For acute and subacute eruptions) These agents act by drying or hardening the skin. Do not use salicylic acid or tannic acid on denuded areas	Alum, U.S.P.	1-5%
	Bismuth Subnitrate, U.S.P.	1-5%
	Boric Acid, U.S.P.	3-15%
	Calcium, U.S.P.	4%
	Kaolin, N.F.	17%
	Salicylic Acid, U.S.P.	10%
	Tannic Acid, N.F.	2-3%
	Zinc Oxide, U.S.P.	1-10%
	Zinc Sulfate, U.S.P.	20-25%
	Ammonium Dichromate, U.S.P.	2-3%
	Althoe, N.F. (for psoriasis)	2-5%
	Potassium Bichromate, U.S.P. (Vioform®)	0.1-0.25%
	Rosin, U.S.P.	3%
KERATOPLASTIC and MILDLY STIMULATING (For subacute and chronic eruptions) These agents produce a counter irritation of the skin, causing inflammatory exudate to subside and redness to be relieved.	Salicylic Acid, U.S.P.	0.25-4%
	Sulfur, Precipitated, U.S.P.	4-6%
	Tannic Acid, U.S.P.	1-6%
	Coal Tar Solution, U.S.P.	3-10%
	[Tar, B.P.] Juniper Tar, U.S.P. (Oil of Cedar, B.P.)	0.5-4%

STIMULATING and/ KERATOLYTIC (For hr i e pti) Th g t t by d e o t ing th h ny l y e r of the k a d by m v g i deb e by de squa m tio Th y als i k n l l y	Ammonit d M Y USP BP	5 10%
	At h l i N F (f p)	0 5 1 0% (C tio)
	Ch y bin USP	0 5 10% (C tio)
	R in l USP, BP	5 10% (Up t 40%)
	S licylic A d, USP, BP	6 10%
	S u f p r e c i p i t d USP, BP	6 10%
	T C l i t e S o l u i d USP	10 20%
	Coal T USP (P e p a r e d C a l t B P) P l e t N F	5 20%
	(T B P) J l p T USP (O u t C a d e B P)	2 10%
	Am m o n i u m M e r c u r y USP BP	2 3%
BACTERICIDAL A i d u f p l e t l i and I f a n i d e s i n o l t m e i s (e t h e r s)	I o d o c h l o h y d y u s USP (V t o m b)	2 3%
	O x y l t y l C h l r i t a c y c l i e o t i a c y l USP	3%
	N e m y u n E y t h m y c i n o r C h l m p h a l i c o l USP	1%
	P o l y m y x i n B S i f t e USP w i t h B i t USP o O y l e t y e l l e USP	3 12%
	B o A d s USP BP	3 10%
FUNGICIDAL	S l i c y l i c A d USP BP	3 10%
	S u f p r e c i p i t d USP BP	3 10%
	Z i n c d i t e,	5 10%
	S o d i m P r p t i N F	5 15% O t m t
	C h l p a c o t a s USP (O u t)	5 10%
PARASITICIDAL	B e n z y l B o a t USP BP	15 30%
	S u l f r p p i a t d USP BP	3 10%
	C n i p h S p t i N F	1 4%
	C o l i T S o l u i d USP	3 15%
	E t h y l A m i n b t e USP BP (B o c a l)	5 10%
ANTIPRURITIC F o r p e r i r i l l f a t b i g A l e y d t m i n e s e f p t u e f l i	H y d o u i USP	1 2 5%
	M t h o l USP, BP	0 2 5 1%
	P h l i USP, BP	0 5 2%
	S l i c y l i c A c i d USP BP	1 2%
	P d p h y l m B e s USP	2 5%
CAUSTICS a d CORROSIVES	S l i y l A c i d P l a t USP	40%

Chapter 6

DISEASES OF THE RESPIRATORY SYSTEM

UPPER RESPIRATORY INFECTIONS

THE COMMON COLD (code No 300 100)

The common cold is a benign inflammation of the mucous membranes of the upper respiratory tract. Part or all of the upper passages may be affected and the manifestation will vary with the areas involved, the severity of the infection and the presence of complications. The etiology has not been determined but a virus is frequently suggested as the possible cause.

The diagnosis is often made by exclusion. Colds must be differentiated from the early stages of many of the communicable diseases which have a similar onset. One is justified in speaking of the common cold or no specific infections only when no specific organism can be found and the disease is primarily localized.

A Local Manifestations. One usually finds inflammation of the mucous membranes of the involved areas.

- 1 Acute Rhinitis (code No 310 100) Nasal congestion and discharge
- 2 Acute Pharyngitis (code No 631 100) Sore throat with pain on swallowing
- 3 Acute Laryngitis (code No 330 100) Hoarseness, pain on swallowing and at times a dry cough

B General Manifestations. Mild generalized aches and pains, sweating and usually a mild fever.

Treatment

No specific treatment is known.

A General Measures.

- 1 Rest. In general, adequate rest is probably completely beneficial. Rest for the first 24-48 hours is of utmost importance in therapy. Patients usually feel much better and the danger of complications is apparently diminished by this regimen.
- 2 Fluid. Patients should be encouraged to drink fluids sufficient to prevent dehydration and to maintain a normal fluid output. There is no evidence that the course of the disease may be influenced in any way by forcing fluids to promote diuresis or by inducing diaphoresis.
- 3 Diet. A palatable, well-balanced diet is to be preferred. Special diets and fasting regimes do not influence the course of the disease.

- 4 Dr gs In gen l t may be stat d th t no drug a e known wh h sh rt the durat n or modify th ev lty of the no pec fic inf tion drugs aim d at preventing compli at s a e f equ ntly of little value

A alg cs nd antipyr tics Th se r often eful in making the p tient omfortabl exc pt fo the diapho esis that is p duc d in a feb ile patient No n compound any bett than anoth r fo this purpo B cau e of the r l w t xicity the s licyl tes are p efe d Aspirin 0 3 0 6 Gm (5 10 gr) very 2 4 hours p n may b given

- b Sulfon m des Th se of s lfom m d s is ment on d only to b ond mn d Th e a e v y few if any i d l at ons fo the f th e drugs in treatment of the c mm n old These drug in n way influ n e th cou e of th d a e m y be toxic and prob bly do l t t l to p ev nt compli at o s Th ppli both to sy tem and local us If complicat ons do d v lop and a am nabl to lfonom des they sho ld th n be g en

- c Antib o t cs Th e ag nts e of no valu in th t eatment of the nsp if infect n Th y may be of real v l in p e ntng ser ompl tion from oc ur ing b t th da gr f nd cng organism res stan e altering the flora f the pl t ryp s s g s r s g d g s n ti at on mak th r r t n use unwis At tim these d ugs m y be us d pr phyl t ally in p tient w th ch onic v l vul h rt d ea to p v nt re u e of heumat c f e s d t p v t ubacute bact l do ard t s f m c ur ing (s e page 172)

- d Antihust min M y b of b f t in r lieving th very a ly sympt ms (e g sne zig and hino rh) There is no id that thes d g abort or alte th course of th mm n old A l ge numb of antih tamin c re mmer ally l bl m y are n w off i lly pt d (U S P o N N D) (s e page 45)

B L l M u L l m d c t ons hav n influ c on th ou s f the d Th y a u d p ma ly to p mot the comf rt f th p ti nt

- i V s onst to Th e mploy d to g ve t mpo y el f f m nas l b truction and/or hin he

a Inhal r (g Amph t min l h lant U S P) may b of b n fit n m l d e

- b N s l s p y r d cps 17 Eph d in S l f t U S P in l n 1/47 Ph nyleph in Hyd ochl rid U S P (N o-Syn ph [®]) in sali e o Mild N ph olin Hyd hlo d S l t i N F (Pr vi [®]) in al e a e t f t y D ot f t n e th v e y 2 3 ho o f p o l g d p od be a v oconst i tor drugs i r t t i g and an s e bound ng tion O l p p t i o s an be s d fo mild mo prol g d s t i t a tion N o Syn ph ine[®] 10 mg (1/6 g) y 4 h r is f f t i n many ase Eph dr n Sulf t U S P Eph dr in Hyd hl d U S P B P 25 50 mg (3/8 3/4 gr) alon mbin d with s b bit t m y ls b mpl yed E c v C N S t mulat on should b id d

- 2 Throat swabs Swabbing the throat with an antiseptic agent is valueless in combating infection or altering the course of the disease and may be harmful. Strong antiseptics may be protein precipitants producing necrotic tissue which can act as a culture medium for pathogenic organisms. Waker solutions are washed away in a matter of minutes.
- 3 Gargling and throat irrigation These are of little value in affecting the disease but the heat of warm non-irritating gargle or irrigation may give marked transient relief of pain in cases of acute pharyngitis. Solution recommended for use are isotonic salt solution (1 tsp salt per quart or liter of water) or 5-20% glucose in water (1-4 tsp glucose or Kao[®] syrup per cup or 240 cc of water).

C Cough Medications The cough associated with an upper respiratory infection is usually caused by dryness and inflammation of the posterior pharynx and upper trachea. As such it represents a physiological protective mechanism against downward drainage of infected material usually requires no therapy and should never be abolished completely. It may be suppressed if it is too exhausting, painful, prevents sleep or interferes with the cause of coexisting conditions (e.g. immunodeficiency). It can be alleviated or suppressed by a number of measures.

- 1 Voluntary suppression of the cough will usually prevent much of the coughing.
- 2 Sugar lozenges (hardops) are usually soothing to the throat.
- 3 Inhalation of warm moist air (steam) usually relieves the cough. Compound Benzoin Tincture U.S.P. 1 tsp may be added to each quart of water but it is the moist air rather than any medication that gives relief.
- 4 Drugs for severe coughs
 - a Codeine Phosphate U.S.P. is the drug of choice. It should be given at an insufficient dosage to suppress but not to abolish cough. Usual dose for the adult is 8-15 mg (1/8-1/4 gr) orally every 4 hours as needed.
 - b Expectorant cough mixture. It is doubtful if any of the expectorant cough mixtures have any effect on increasing the bronchial secretion or on altering the viscosity of the mucus. Some of the syrupy cough mixtures have a temporary soothing action in the oropharynx but bitter and more prolonged side effects are obtained with sweetened lozenges (cough drops). The cough mixtures include Terpin Hydrate Elixir N.F. syrup of ammonium hydrochloride, Tolu Balsam Syrup U.S.P. Wild Cherry Syrup U.S.P. etc. The action of codeine is not enhanced in any way when it is added to the mixtures and it can now be given in tablet form.

D Local Heat Marked relief in the nose and throat can often be obtained from steam inhalations or exposure to warmth (hot water bottle or infra-red lamp over the nasal region).

Treatment of Complications

The principal complications of the common cold are extension to accessory structures in direct contact with the upper respiratory passages or secondary bacterial invasion of the mucous membranes.

and accessory structures. These usually require antibiotics or sulfonamides and may require the attention of an otolaryngologist.

Prophylaxis

The principal prophylaxis is similar to that of any contagious disease: avoidance of exposure whenever possible; avoidance of sudden changes of temperature and excessive fatigue. Administration of large doses of any of the vitamins, cold vaccines orally or by injection, gamma globulin or 'hardening up' have all proved valueless in preventing or in altering the course of the disease.

ACUTE SINUSITIS (code No 32.130)

The acute infection of the paranasal sinuses following upper respiratory infections is usually caused by secondary bacterial invasion. The infecting organisms most frequently are streptococci, staphylococci or pneumococci.

Treatment

A. Specific Measures

1. Penicillin is the drug of choice since most of the organisms are penicillin sensitive. It is administered as follows: 300,000 units penicillin procaine I.M. once or twice daily. Other wide spectrum antibiotics may also be employed.
2. Local administration of penicillin and other antibiotics by nose drops and use of negative pressure is still difficult to evaluate.

B. General Measures

1. Bed rest.
2. Local external heat over the sinuses.
3. Analgesic. Aspirin or codeine may be used.
4. Vasoconstrictor drugs. Non-irritating nose drops may be used to facilitate drainage; or drug may be given in tablet form by mouth for similar effect (see page 109).

C. Duration of treatment in sinusitis during acute sinusitis

EPISTAXIS (code No 301)

Epistaxis may be due to a variety of diseases or disorders.

A. Predisposing Factors Blood dyscrasias, hypertension, arteriole sclerosis, prothrombin deficiency (e.g. cirrhosis of the liver), nasal ulceration, nasal angioma and retention of the infectious discharges (e.g. measles and diphtheria).

B. Predisposing Factors External trauma to the nose, violent blowing of the nose, sneezing, picking of the nose, increase of existing high blood pressure or lowering of atmospheric pressure.

C. Location The bleeding is most frequently on the anterior portion of the nasal septum, less often the end of the inferior and middle turbinates and rarely further posteriorly.

2 Throat swabs Swabbing the throat with an antiseptic agent is valueless in combating infection or altering the course of the disease and may be harmful. Strong antiseptics may be protein precipitants producing necrotic tissue which can act as a culture medium for pathogenic organisms. Weak solutions are washed away in a matter of minutes.

3 Gargling and throat irrigation. There is of little value in affecting the disease but the heat of a warm non-irritating gargle or irrigation may give marked transient relief especially in cases of acute pharyngitis. Solutions recommended for use are isotonic salt solution (1 tsp salt per quart of water) or 5-20% glucose in water (1-4 tsp glucose or Karo® syrup per cup of 240 cc of water).

C Cough Medications. The cough associated with an upper respiratory infection is usually caused by dryness and inflammation of the posterior pharynx and upper trachea. As such it represents a physiological protective mechanism against downward drainage of infected material. Usually requires no therapy and should never be abolished completely. It may be suppressed if it is too exhausting, painful, prevents sleep or is otherwise a direct cause of coexisting conditions (e.g. immediately postoperative). It can be alleviated or suppressed by a number of measures.

1 Voluntary suppression of the cough will usually prevent much of the coughing.

2 Sugar lozenges (cough drops) are usually soothing to the throat.

3 Inhalation of warm moist air (steam) is usually very soothing. Compound Benzoin Tincture U.S.P. 1 tsp may be added to each quart of water but it is the moist rather than any medicinal action that gives relief.

4 Drugs for severe coughs.

a Codeine Phosphate U.S.P. is the drug of choice. It should be given alone in sufficient dose to suppress but not to abolish cough. Usual dose for this is 8-15 mg ($\frac{1}{8}$ - $\frac{1}{4}$ gr) orally every 2-4 hours per os.

b Expectorant cough mixtures. It is doubtful if any of the expectorant cough medicines have any effect on increasing the bronchial secretion or altering the viscosity of the mucus. Some of the syrupy cough mixtures have a temporary soothing action on the oropharynx but bitter and more prolonged relief are obtained with sweetened lozenges (cough drops). The cough mixture includes Terpin Hydrate Elixir N.F. syrup of ammonium chloride Tolu Balsam Syrup U.S.P. Wild Cherry Syrup U.S.P. etc. The action of codeine is not enhanced in any way when it is added to the mixtures and it can as well be given in tablet form.

D Local Heat. Marked relief in the nose and throat can often be obtained from steam inhalations or exposure to warmth (hot water bottle or infra-red lamp over the nasal region).

Treatment of Complications

The principal complications of the common cold are extension to accessory sinuses in direct contact with the upper respiratory passages or secondary bacterial invasion of the mucous membranes.

and accessory structures. These usually require antibiotics or sulfonamides and may require the attention of an otolaryngologist.

Prophylaxis

The principal prophylaxis is similar to that of any contagious disease: avoidance of exposure whenever possible; avoidance of sudden changes of temperature and excessive fatigue. Administration of large doses of any of the vitamins, cold vaccines orally or by injection, gamma globulin or hardening up have all proved valueless in preventing or in altering the course of the disease.

ACUTE SINUSITIS (code No. 32.130)

The acute infection of the paranasal sinuses following upper respiratory infections is usually caused by secondary bacterial invasion. The infecting organisms most frequently are streptococci, staphylococci, or pneumococci.

Treatment

A. Specific Measures

1. Penicillin is the drug of choice since most of the organisms are penicillin sensitive. It is administered as follows: 300,000 units penicillin procaine I.M. once or twice daily. Other wide-spectrum antibiotics may also be employed.
2. Local administration of penicillin and other antibiotic by nose drops and use of negative pressure is still difficult to evaluate.

B. General Measures

1. Bed rest.
2. Local external heat over the sinus.
3. Analgesic: Aspirin or codeine may be used.
4. Vasoconstrictor drugs: Non-irritating nose drops may be used to facilitate drainage; or drug may be given in tablet form by mouth for systemic effect (see page 109).

C. Do not instill anything in sinuses during acute sinusitis

EPISTAXIS (code No. 301)

Epistaxis or nose bleeding may be due to a variety of diseases or disorders.

- #### A. Predisposing Factors
- Blood dyscrasias: hypertension, arteriosclerosis, prothrombin deficiency (e.g., liver disease of the liver), nasal ulceration, nasal angioma, and certain of the infectious diseases (e.g., measles and rheumat fever).

- #### B. Predisposing Factor
- External trauma to the nose: violent blowing of the nose; picking of nose; increase of existing high blood pressure; lowering of atmospheric pressure.

- #### C. Location
- The bleeding site is most frequently on the anterior portion of the nasal septum, less often at the end of the inferior and middle turbinates and rarely further posteriorly.

- 2 Throat swabs Swabbing the throat with an antiseptic agent is valueless in combating infection or altering the course of the disease and may be harmful. Strong antiseptics may be protein precipitants producing necrotic tissue which can act as a culture medium for pathogenic organisms. Weak solutions are washed away in a matter of minutes.
- 3 Gargling and throat irrigation These are of little value in affecting the disease but the heat of warm non-irritating gargle or irrigation may give marked transient relief of pain in cases of acute pharyngitis. Solutions recommended for use are isotonic salt solution (1 tsp salt per quart or liter of water) or 5-20% glucose in water (1-4 tsp glucose or Karo® syrup per cup or 240 cc of water).

C Cough Medication The cough associated with an upper respiratory infection is usually caused by dryness and inflammation of the posterior pharynx and upper trachea. A cough reflex is a physiological protective mechanism against downward drainage of infected material usually requires no therapy and should never be abolished completely. It may be suppressed if it is too exhausting, painful, prevents sleep or is contraindicated because of existing conditions (e.g. immediately postoperative). It can be alleviated or suppressed by a number of measures:

- 1 Voluntary suppression of the cough will usually prevent much of the coughing.
- 2 Sugar lozenges (cough drops) are usually soothing to the throat.
- 3 Inhalation of warm moist air (steam) is usually very soothing. Compound Benzoin Tincture U.S.P. 1 tsp may be added to each quart of water but it is the most irritating than any medication that gives relief.
- 4 Drug for severe coughs

Codeine Phosphate U.S.P. is the drug of choice. It should be given alone in sufficient dosage to suppress but not to abolish cough. Usual dose for the adult is 8-15 mg (1/8-1/4 gr) orally every 2-4 hours as needed.

- b Expectorant cough mixtures It is doubtful if any of the expectorant cough medicines have any effect on increasing the bronchial secretion although the viscosity of the mucus. Some of the syrupy cough mixtures have a temporary soothing action on the oropharynx but do not and more prolonged results are obtained with sweetened lozenges (cough drops). The cough mixtures include Terpin Hydrate Elixir N.F. syrup of ammonium chloride Tolu Balsam Syrup U.S.P. Wild Cherry Syrup U.S.P. etc. The action of codeine is not enhanced in any way when it is added to the mixture and it can always be given in tablet form.

D Local Heat Marked relief in the nose and throat can often be obtained from steam inhalations or exposure to warmth (as hot water bottle or infrared lamp over the nasal region).

Treatment of Complications

The principal complications of the common cold are extension to accessory structures in direct contact with the upper respiratory passages or secondary bacterial invasion of the mucous membranes.

that the maximum desensitization may be produced

1. Determine the offending allergen. This is usually accomplished by a careful history plus skin tests with antigens of known allergenic activity
2. Course of injection of allergen extracts. The material is given subcutaneously weekly or biweekly in increasing dosages. An average schedule that requires 10-20 weeks for desensitization is shown below. (This course must be repeated every year.)

Week	Dilution	Size of Dose	Noon or Pollen Units	Week	Dilution	Size of Dose	Noon or Pollen Units
1	1:5000	0.1 cc	20	11	1:50	0.1 cc	2000
2		0.2 cc	40	12		0.2 cc	4000
3		0.3 cc	60	13		0.3 cc	6000
4		0.4 cc	80	14		0.4 cc	8000
5		0.5 cc	100	15		0.5 cc	10,000
6	1:500	0.1 cc	200	16		0.6 cc	12,000
7		0.2 cc	400	17		0.7 cc	14,000
8		0.3 cc	600	18		0.8 cc	16,000
9		0.4 cc	800	19		0.9 cc	18,000
10		0.5 cc	1000	20		1.0 cc	20,000

If a shorter schedule must be employed, one can cut the above schedule in half by eliminating the 0.2 and 0.4 cc doses in schedules. However, in so doing, utility must be exercised to avoid reaction.

3. Maintenance dosages. When patient has completed the program of desensitization and hay fever season has begun, administer 0.2 cc of 1:50 dilution of extract (4000 noon units) every 1-2 weeks until season is over.
4. Continuous desensitization. If patient receives 0.2 cc of 1:50 dilution every 2 weeks throughout the year, the annual "boost" course can be started later, beginning with 0.1 cc of a 1:50 dilution (day 11) and continuing with above or a selected schedule.
5. Attempt at desensitization on hay fever has begun generally met with little success.

B. General Measures

1. Drug

Cortisone (ACTH) and the corticosteroids have been shown to give complete relief from allergic rhinitis within 12-36 hours. If the offending allergen is apparent within the day is withdrawn, symptoms usually return but only individual with seasonal hay fever may be latently free of the duration of the hay fever season.

- a. Antihistaminic drugs give relief in 60-80% of patients initially; however, the relief tends to wear off as the season continues. Many antihistaminics are available, all have similar side effects of lethargy and drowsiness. The dose is variable with some preparations (page 45).

- c. Ephedrine sulfate or ephedrine hydrochloride 25-50 mg

Treatment

- A Local Measures Have patient sit erect with head forward. If reclining there is danger of aspiration of blood.
- 1 Pressure over the bleeding site is usually all that is necessary. A small pledget of cotton moistened with hydrogen peroxide will usually stop the bleeding.
 - 2 Cauterization. When active bleeding has ceased touching the bleeding point with a bead of chromium trioxide (chromic acid) or trichloroacetic acid will usually prevent further bleeding. Electrocautery is also satisfactory.
 - 3 Severe bleeding in the anterior part of the nose can usually be controlled with a tampon introduced through the nostril. If bleeding is posterior it may be necessary to introduce a posterior nasal pack. This is done by the use of two strings attaching one string near each end of a rolled 2 x 2 or 4 x 4 gauze pad. A third string is tied at the middle of the rolled pad. A soft catheter is then introduced into the nasopharynx through one nostril and pulled out through the mouth. The end of one of the first two strings is tied to the oral portion of the catheter and pulled back through the nostril. The string is then removed from the catheter and the procedure repeated pulling the second string through the other nostril. The pack is guided through the mouth into the nasopharynx and pulled into place by traction on both strings. These are then tied over pad under the nasal septum. The third string is taped to the face and used later to remove the pack. Do not leave pack in more than 48 hours.
 - 4 Ice packs on the nose or to the back of neck are of no benefit.
- B Specific Measure Treat the underlying disease.

ACUTE TONSILLITIS (code No 634 100)

Acute tonsillitis is an infection of the faucial tonsils caused by any of a number of organisms. It is characterized by both local and generalized symptoms of varying degree.

Treatment

Depends on the causal organism. (For streptococcus see page 481.)

ALLERGIC RHINITIS (Hay Fever) (code No 310 392)

Hay fever is an allergic disorder which usually occurs in spring or summer and is characterized by rhinitis, sneezing, nasal obstruction, redness of eyes, and itching of the nose, throat, or eyes. Pollens are the most common allergens.

Treatment

A Specific Measures There is no true specific treatment. Hypo-sensitization or desensitization is frequently beneficial and consists of administration of gradually increasing doses of the allergen (usually pollen) so as to induce an immunity in the susceptible individual. For best results therapy should be started 3-6 months before the onset of the hay fever season so

- 3 Sleepiness Pentobarbital Sodium U S P Pentobarbital Sodium B P 0.1 Gm (1½ gr) at bedtime should be given

CHRONIC TRACHEOBRONCHITIS (Chronic Bronchitis code No 350 100 0)

A chronic nonspecific inflammation of the tracheobronchial tree manifested by cough which is usually the only constant symptom. The cough may be productive or nonproductive. *Do not diagnose chronic bronchitis on the basis of chronic cough alone.* Any patient with a chronic cough should be given a thorough examination including a chest x-ray. As a rule a diagnosis of chronic bronchitis can be made only by exclusion. Primary chronic bronchitis is a rare disease. Almost all cases of chronic bronchitis are secondary to other respiratory conditions or inflammations. Intratable cases may have an allergic basis. Physical findings may be absent or a few rhonchi and wheezes may be heard.

Treatment

A Specific Measures There is no specific treatment for chronic bronchitis. *Treat the underlying condition.*

B General Measures

- 1 Remove or eliminate exciting causes such as smoking, exposure to cold, air damp atmosphere, industrial fumes, etc.
- 2 Drug: Ephedrine and similar drug given by mouth or nebulization e.g. Isoproterenol Hydrochloride U S P (Aludrine® Isuprel®) gives relief in many cases where bronchospasm is present. Potassium Iodide Solution N F 5-10 drops tid as tolerated may be helpful.
- 3 Adequate rest in dust-free room.
- 4 Optimum nutrition and hygiene.

BRONCHIAL ASTHMA (code No 350 390)

Bronchial asthma is a symptom complex due to a variety of

It is characterized by dyspnea especially of the expiratory type with wheezing and whistling which are due to edema of the bronchial lining and/or constriction of the smooth muscle leading to constriction of the bronchi.

Diagnosis

A History of paroxysmal attacks of expiratory dyspnea frequently in an otherwise healthy patient with a definite allergic tendency. The attacks are precipitated by exposure to the allergen or triggered by various motives.

B Physical Examination

- 1 During typical attack Examination characteristic:
 - a Severe expiratory dyspnea and asthmatic cyanosis
 - b Chest held in partial inspiratory position
 - c Inspiration short and expiration greatly prolonged
 - d Cough difficult and may become violent
 - e Sputum thick and tenacious
 - f Chest hyperresonant to percussion

- with a barbiturate every 4 to 6 hours may give relief (see page 109)
- d Sedation may be of value if patient is nervous or upset (see page 39)
 - 2 Allergen free atmosphere
 - a Dust proof respirator masks can be used during the hay fever season
 - b Air conditioning equipment to filter the air entering the patient's room may prove valuable
 - c An area free of the offending pollen can be visited during the pollinating period
 - d When dust is the offending agent prepare a dust free bed room as follows. Cover mattress and pillow with an allergen free material (plastic or sheet rubber). Remove all rugs carpets drapes bedspreads or other lint producing materials and remove all ornate furniture or other objects which are not easily dusted
 - e Household pets must be considered possible sources of allergens

DISEASES OF THE BRONCHI

ACUTE TRACHEOBRONCHITIS

(Tracheitis code No 340 100) (Bronchitis code No 350 100)

An acute nonspecific inflammation of the trachea bronchial tree usually following an acute rhinitis or pharyngitis and usually accompanied by a low grade fever. A productive or nonproductive cough is present. Physical examination may be entirely negative or scattered coarse rhonchi may be heard over the chest.

Treatment

A Specific Measures

- 1 There is no specific treatment unless secondary infection is present
- 2 In severe cases prescribe inhalation by aerosol inhalation (see page 153) several times daily and/or penicillin procaine 300 000 units I.M. once or twice daily may be of value

B General Measures

- 1 Rest. Bed rest is most important in shortening the course of the disease
 - 2 Control of cough
 - a Steam inhalation or eucalyptus water or saline. The hacking cough of the dry stage is helped best by steam inhalations in a warm room. Compound benzoin tincture 1 tsp to each qt (liter) of water may be added
 - b Codeine in small doses 15-30 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr) q 3-4 hours may be given to help control the paroxysms of coughing
 - c Ephedrine cough mixture will be helpful if bronchospasm is present (manifested by wheezing)
 - d Ephedrine sulfate or

hydrochloride 1% Sol	30 0 3i
Codeine phosphat	0 5 gr viiss
Syrupy vehicle q s ad	120 0 iv
- Sig 4cc (1 tsp) q 3-4 hours p r n cough

B. S. e. Att. k. Epinephrine responsive patients (many also follow for treatment asthma table below)

1. Epinephrine injection (mg) 1 cc of adrenaline 0.5 1 0 cc (3 15 mg) 1000 solution about and repeat every 30 60 minutes if necessary
2. Epinephrine inhalation no isoproterenol (isoprel[®]) 1 100 by ebullient oxygen for spray may give dramatic relief. May repeat every 30 60 minutes
3. Epinephrine in oil 1 500 0.2 1 0 cc (3 15 mg) I.M. for prolonged effect. May help prevent recurrence of attack and can be repeated in 10 14 hours
4. Aminophylline (theophylline ethylenediamine) 0.24 0.48 Gm (3 3/4 7 1/2 gr) in 10 20 cc (2 1/2 5 dr) saline slowly I.V. if attack not controlled. May also give this as rectal instillation or rectal suppository in the same dose
5. Sedation must be adequate. Use one of the following:
 - a. Pentobarbital sodium (pentobarbital sodium) 0.1 Gm (1 1/2 gr) at once and may repeat
 - b. Paraldehyde 4 8 (1 2 dr) orally in fruit juice or rectally in 30 cc (1) oil
6. 100% oxygen (or 20% helium with 80% oxygen) inhalation by mask at 6 12 ltr/minute may give great relief from dyspnea
7. When available the use of oxygen by intermittent positive pressure (e.g. Bunnitt valve) and bronchodilating aerosols administered simultaneously through the same apparatus affords the most dramatic relief in acute attack of asthma. As a bronchodilator isoproterenol (isoprel[®]) to be preferred because it produces a lesser degree of systemic effect than diphenhydramine
8. The plasma fluid retention lowering agents (e.g. Alveol[®]) depolymerizing nym (e.g. hyaluronidase) or digoxin nym (e.g. typhoid) by osmosis in this condition is still not definitely advised. The advisability of using the latter has recently been questioned in view of its effects on the morphology of the cells of the capillary lining as well as its irritating local effect

C. Status Asthmaticus and Severe Attacks in Epinephrine-resistant

Patient

1. Corticotropin (ACTH) 25 50 units of gel preparation subcutaneous I.M. or intravenous or orally immediately and repeat every 6 hours. If the patient hospitalized administer corticotropin (ACTH) 20 40 mg I.V. drip over 6 12 hours or orally every 24 hours. ACTH may have a morphine-like action but otherwise both are about equally effective. Relief should be evident in 6 12 hours and almost complete relief should be obtained with all manifestations in 24 48 hours. The medication should probably be continued for 7 10 days in gradually diminishing doses after the first 4 5 days
2. Patient should be hospitalized if possible in an allergen-free room
3. 100% oxygen or 20% helium with 80% oxygen should be given by mask for relief of dyspnea
4. Aminophylline (theophylline ethylenediamine) 0.24 0.48 Gm

g Chest full of musical rales which frequently may be heard at a distance

2 Between attacks May be entirely negative or show only scattered expiratory wheezes

C Differential Diagnosis Many diseases may simulate bronchial asthma. Of these the most important are cardiac asthma and generalized emphysema especially when chronic bronchitis is superimposed. The appearance of bronchial asthma in middle or old age should make one suspect bronchial neoplasm.

Treatment

A The treatment may be divided into two phases

1 Treatment of the actual attack

2 Interim therapy which is aimed at preventing further attacks

B Drugs used for the specific treatment of asthma. In the control of the acute attack certain therapeutic agents have come to be looked upon as specific for relief while other preparations are of value in aborting attacks. It is necessary to know the preparations available, their mode of administration and the indications for each. The chart on page 118 summarizes the important drugs.

Epinephrine (drenaline) is the drug of choice for the emergency management of acute bronchial asthma. However it has been shown that corticotropin (ACTH) or cortisone can be used to stop an attack when all other measures fail. The onset of action of ACTH and cortisone is much slower than that of epinephrine but they should be employed concurrently with epinephrine in severe attacks of asthma. *Epinephrine must be used cautiously in patients with cardiac asthma, hypertension or angina.*

Treatment of the Acute Attack. Do not use morphine

A Mild or Moderate Attack Epinephrine (drenaline) is the drug of choice

1 Epinephrine injection (injection of adrenalin) 0.2 to 0.5 cc (3 to 8 min) of 1:1000 bicut

2 Epinephrine inhalation (1:100 in aqueous solution) by nebulizer every 30 to 60 minutes per nebulizer (see page 118)

3 Moderate attack Repeat epinephrine (drenaline) subcutaneous every 1 to 2 hours

4 Epinephrine nebulizer 0.2 to 1.0 cc (3 to 15 min) 1:500 I.M. may also be given at onset if a prolonged effect is desired. May repeat in 10 to 14 hours

5 Aminophylline (theophylline ethylenediamine) 0.24 to 0.48 Gm (3 3/4 to 7 1/2 gr) in 10 to 20 cc (2 1/2 to 5 dr) saline 1 way I.V. If still not relieved 0.48 Gm (7 1/2 gr) may be added to 500 to 1000 cc of saline and given by I.V. drip. May also give this as rectal instillation or rectal suppository in the same dose

6 Ephedrine sulfate or hydrochloride 25 to 50 mg (3/8 to 3/4 gr) with or without a barbiturate may relieve mild attack (see page 118)

7 Reassure patient that attacks can be controlled

8 Sedation Phenobarbital (Phenobarbital) 0.1 Gm (1 1/2 gr) immediately may repeat 0.03 Gm (1/2 gr) q.i.d.

(3 3/4 7 1/2 gr) in 10 20 cc (2 1/2 5 dr) saline slowly I V and by rectal suppositories for immediate relief of symptoms 0.48 Gm (7 1/2 gr) may be added to 500 1000 cc of saline and given by I V drip

5 Sedation must be deferred until relief is obtained Use one of the following

a Pentobarbital sodium (pentobarbitone sodium) 0.1 0.2 Gm (1 1/2 3 gr)

b Paraldehyde 8 15 (2 4 dr) in 30 cc (1 oz) oil by rectum

6 Surface tension lowering agents (Alvirene[®]) by nebulizer may be helpful in some cases (see page 153)

7 If corticotropin (ACTH) or corticosteroids are not available

a As soon as epinephrine responsiveness returns use epinephrine as above Epinephrine may be administered cautiously i.e. 1:1000 solution in 1 liter of 5% glucose by intravenous drip (50 80 drops per minute)

b General anesthetic agents may be life saving

(1) Rapid instillation of 30 90 cc (1 3 oz) of ether in equal quantities of oil oil repeat in 12 to 24 hours if necessary Usually patient wakes free of attack

(2) If available anesthetic available inhalation through the mask may be employed

8 Bronchocopy under general anesthesia is sometimes indicated to move intrusions

D General Measures

1 Eliminate any known allergen if compatible with environment

2 Maintain adequate rest and relief of hyperinflation by reassurance and diet

3 Respiratory infections must be treated vigorously with antibiotics directed I M or by aerosol

4 Fluids orally or parenterally to prevent dehydration

Intermittent Therapy

A Specific Therapy Attempt to determine which all agent plays a role and treat accordingly

B General Measures

1 Emotional disturbance should be corrected whenever possible

2 Good living hygiene should be promoted

3 Patient with apparently intermittent asthma (usually due to infection of bronchi) may be helped by antibiotic therapy (see page 114 and 120)

4 Ephedrine hydrochloride 25 50 mg (3/8 3/4 gr) with or without phenobarbital (phenobarbital) 15 30 mg (1/4 1/2 gr) every 3 6 hours may prevent or delay occurrence

5 Aminophylline phenethylphenobarbital pulses

 & Aminophylline 0.2 g iii

 Ephedrine hydrochloride sulfate 0.025 g 3/8

 Phenobarbital 0.015 gr 1/4

Sig 1 pulse every 4 hours

6 Antihistaminic agents may give relief in some patients but this is usually bronchial asthma generally bronchial asthma diagnosis appointing (see page 45)

DRUGS USED IN THE TREATMENT OF BRONCHIAL ASTHMA

Preparation	Dose	Mode of Administration and Indication
Epinephrine Injection U S P Injection of Adrenaline B P (1:1000 dilution of the hydrochloride in aqueous solution)	0.2 to 0.5 cc (3 to 15 min) repeat up to q 30 minutes if necessary 1 cc (15 min) in liter of 5% glucose solution Give at 60 to 80 drops per minute	Subcut This is the most commonly used preparation I V Caution Reserved for very severe acute attacks when more conservative measures fail
Epinephrine in Oil Injection U S P (1:500 dilution)	0.2 to 0.5 cc (3 to 15 min) May repeat in 10 to 14 hours Duration of action is 3 to 4 hours	Subcut or I M usually given with aqueous epinephrine to patients with severe or current asthma
Epinephrine Inhalation U S P (1:100 dilution in aqueous solution)	0.5 cc in nebulizer Individualize dose 4 to 8 inhalations usually suffice	Glass nebulizer operated by hand bulb or pressure from an oxygen tank or nebulizer with intermittent positive pressure breathing (IPPB) (see page 148) Most useful in aborting attacks
Isoproterenol U S P (Isuprel®) (1:100 dilution in aqueous solution) For inhalation only	(Isoproterenol causes less vasoconstriction)	
Isoproterenol tablets (5, 10, 15 mg)	Sublingual, individualized	May be useful in aborting attack
Corticotropin (ACTH)	10 to 40 mg by continuous I V drip over 24 hours or 25 mg of regular or gel I M every 6 hours initially	Decreases tolerance in prolonged use Used for severe attacks and status asthmaticus
Cortisone	See page 423	Orally
Aminophylline Injection U S P B P (Theophylline Ethylenediamine)	0.24 to 0.48 Gm (3/4 to 7/8 gr) in 10 or 20 cc saline May repeat in 3 to 4 hours Duration of action 1 to 3 hours	I V slowly May be used with or without epinephrine Valuable in severe attacks when patient is epinephrine fast
Aminophylline U S P rectal suppository	1 suppository every 12 hours	Useful only when prolonged life is desired
Ephedrine Hydrochloride U S P B P or Ephedrine Sulfate U S P (capsule or pill)	25 to 50 mg (3/8 to 3/4 gr) every 3 to 6 hours May combine with phenobarbital 15 to 30 mg (1/4 to 1/2 gr) or phenobarbital sodium 30 mg (1/2 gr)	Orally Of little use in acute attack May be of some value in aborting an attack or decreasing the number of attacks
Antihistamine drugs	See page 45 for dosage	Orally Of little use in acute attacks May be of value in aborting attack or decreasing number of attacks

bronchitis appears to be of less value than direct inhalations

- 3 Streptomycin in aerosol (see page 153) may also be of benefit in some patients especially those in whom penicillin resistance exists. Each cc should contain 50-250 mg of streptomycin depending on concentration desired. Administer in the same manner as for penicillin (see above)
- 4 Combined penicillin-streptomycin in aerosol may be of benefit in many cases. Use the same concentration for the drugs as used individually
- 5 Oxetyazoline (T mycin®) is also available for aerosol administration (50 mg/cc in propylene glycol) and may be used as above (See page 153)
- 6 Erythromycin may be of value in obtaining better drainage of thick mucus in patients (see page 153)

B. General Measures

- 1 Postural drainage. Postural drainage has proved to be the most effective angle measure for the symptomatic relief of patients with bronchiectasis. The patient should assume the position that gives him the maximum drainage and this varies with the location of the lesion. Experience will help the patient determine the best position to assume. Since most lesions are at the lung bases the most common method is for the patient to lie prone across the edge of the bed with folded arms resting on a pillow on the floor maintaining this position for 10-15 minutes. Two to four times a day is usually sufficient. The first drainage being just upon awakening and the last just before bedtime.
- 2 Avoidance of upper respiratory infection is very important in controlling the bronchial infection.
- 3 Correction of acidities. Many patients with bronchiectasis suffer from chronic upper respiratory infections with postnasal drip. This must be corrected whenever possible.
- 4 Climate. Although climate does not usually make a difference in the onset of bronchiectasis especially in the winter due to the incidence of upper respiratory infections. Avoid a dusty or smoke-filled atmosphere.
- 5 Rest. Patient with severe disease should always have adequate rest in bed for symptoms often ameliorated by this measure. The foot of the bed should be raised 6 to 12 inches.
- 6 Good nutrition and health are very important. Adequate food and rest will aid in lowering the progress of the disease. Smoking should be prohibited.
- 7 Bronchoscopic drainage is of value initially in all cases to eliminate bronchial tenacious obstruction containing factors which may be necessary to dilate the tenacious bronchus but repeated bronchoscopies are not advised.

C. Surgical Treatment Presently accepted indication in lung

- 1 Young patients in good condition who are having chronic recurring symptoms of any degree. Modern surgery will permit resection of fairly extensive bilateral disease.
- 2 Patients up to 60 years of age who are having severe symptoms

- 7 Patients who are not helped by other measures may be treated chronically with small doses of corticotropin (ACTH) or one of the cortisones. The dosage employed is just sufficient to keep them comfortable and relatively free of symptoms.

BRONCHIECTASIS (code No 350 100 6)

Bronchiectasis is a chronic progressive disease of the bronchi and bronchioles characterized by dilatation of the bronchi or bronchioles, the presence of varying amounts of infiltrative infection and finally destruction of the involved parts and of the surrounding tissue. The etiology in many cases is unknown but congenital factors and chronic or recurrent pulmonary infections undoubtedly play a role.

Diagnosis

A History

- 1 Chronic cough usually productive of much purulent sputum and more marked upon arising in the morning.
- 2 Recurrent attacks of pulmonary infections with aggravated cough, fever, sweats and chills. Hemoptysis is common.

B Physical Examination Chest findings except in severe cases or during acute pneumonitis are rarely significant. Rales at lung bases and some wheezing or pectoriloquy are the most common findings.

C Laboratory Findings

- 1 X-ray Routine chest x-rays are usually insufficient to make diagnosis. If chest x-ray is negative and bronchiectasis is suspected further studies are necessary.
 - a Bronchoscopic examination
 - b Bronchograms (x-rays of chest following instillation of iodized oil into bronchi either through bronchoscope or directly into the trachea) are most important for diagnosis. These must be made by an experienced radiologist.
- 2 Sputum The sputum is usually found to separate into 3 layers. Bacteriological studies always reveal mixed infections usually with streptococci and staphylococci predominating.

Treatment

A Specific Measures Treatment with antibiotic agents has been of benefit in temporarily ameliorating symptoms especially during the acute exacerbations but has had no lasting effect. The amount of sputum and cough is reduced and the patient feels better but these benefits wear off in a few weeks to months. The treatment should be repeated as necessary and may be of greater benefit in milder cases or in debilitated patients where surgery is considered too radical a procedure. When possible predominant organisms should be identified and their sensitivity to the various antibiotics determined.

- 1 Penicillinase resistant (see page 153 for technique) 50,000 to 100,000 units of crystalline penicillin G in 10 to 20 cc saline solution q.i.d.
- 2 Parenteral use of penicillin except during attacks of acute

A Mild to Moderate Cases Penicillin is the drug of choice although broad spectrum antibiotics or sulfonamides are almost equally effective. Equal mixtures of sulfadiazine and sulfamerazine or sulfisoxazole (Gantisin®) are to be preferred whenever sulfonamide drugs are demonstrated (see page 486).

1 Penicillin

a Dosage

- (1) Slow absorption type Penicillin procaine in aqueous suspension or in oil 300 000 units I.M. daily will usually suffice. 300 000 unit bid may be desirable in some cases.
- (2) Intermediate I.M. penicillin 15 000 to 20 000 unit aqueous penicillin very 3-4 hours.
- (3) Intermediate I.M. penicillin 100 000 units aqueous penicillin very 8 hours.
- (4) Penicillin 200 000 units by mouth every 4 hours is usually satisfactory but generally open penicillin should be reserved for use after initial favorable response to parenteral penicillin.

b Duration Continue penicillin until patient has been afebrile for at least 72 hours and white blood count normal.

2 Broad spectrum antibiotics Continue until patient has been afebrile for 48-72 hours and white blood count is normal.

Chlortetracycline Hydrochloride U.S.P. (Achromycin®) 0.25 Gm q 6 h.

b Oxytetracycline Hydrochloride U.S.P. (Tetracycline®) 0.5 Gm q 6 hours.

c Tetracycline U.S.P. 0.25 Gm q 6 h.

d Chloramphenicol U.S.P. (Chloromycetin®) 0.5 Gm q 6 h u.s.

Erythromycin U.S.P. (Erythrocin®) 0.3-0.5 Gm q 6 h r.

3 Sulfonamides

a Dosage (Give 2 Gm of m-bica-bo-te with a h Gm of sulfanamide.)

(1) Sulfadiazine U.S.P. 4-5 Gm (60-75 g) at once orally and 1 Gm (15 g) very 4 hours.

(2) Sulfadiazine and Sulfamerazine U.S.P. 2-2.5 Gm (30-37½ gr) of each orally and 0.5 Gm (7½ g) of each every 4 hours.

(3) Sulfisoxazole U.S.P. (Gantisin®) 4-5 Gm (60-75 g) orally at once and 1 Gm (15 gr) very 4 hours.

b Duration Continue therapy until patient has been afebrile for 48-72 hours and white blood count is normal.

B Moderate to Severe Cases Penicillin to be preferred but broad spectrum antibiotics or sulfonamides may be used in patients intolerant to penicillin.

1 Penicillin In severe cases give intermediate I.M. aqueous penicillin in doses of 100 000 unit very 3 hours day and night or 1 million units of penicillin procaine I.M. very 12 hours. Continue penicillin therapy until patient has been afebrile for 72 hours and white blood count is normal.

2 Broad spectrum antibiotics. Onset of tetracycline 0.5 Gm very 6 h.

(especially recurrent hemorrhage) from predominantly unilateral disease and who are otherwise good surgical risks

DISEASES OF THE LUNGS

PNEUMONIAS

Pneumonia consists of inflammatory changes of the parenchyma of the lung almost always associated with or due to infection. In the past it was customary to classify the pneumonias into anatomic types (i.e., lobar and bronchial) but this classification serves little useful purpose at present. In view of the ever growing number of antimicrobial agents one should classify the pneumonias on the basis of etiology and treat with the appropriate agent.

The pneumonias are still the most common cause of pneumonia. In general it may be stated that the management of all the pneumonias follows the same principles as those governing the management of pneumococcal pneumonia.

PNEUMOCOCCIC PNEUMONIA (code No. 350.101)

Diagnosis

- A. History Usually sudden onset with chills and fever. Often pleuritic pain is present with cough.
- B. Physical Finding Vary with extent of infection and duration of the process from nonproductive rales to massive consolidation of one or more lobes.
- C. Laboratory
 1. Sputum Purulent usually tinged with blood (light pink to the color of prune juice).
 2. All patients with pneumonia (especially if severe) should have the following laboratory examinations in addition to routine urine and blood studies: chest x-ray, sputum smear and blood culture.

Treatment

Before beginning therapy it is advisable to obtain sputum and blood for culture in order to determine the exact bacterial invader. This is important if the infection is severe.

Therapy varies with the severity of the disease depending on the criteria shown in the chart on page 125 and treat as outlined below.

Specific Measures

Penicillin is the drug of choice in pneumococcal infections. Chlorotetracycline, oxytetracycline, tetracycline, chloramphenicol and erythromycin are also highly effective in most pneumococcal infections but are probably slightly inferior to penicillin in severe infections. The sulfoamide drugs are also effective but their response to penicillin is usually more rapid and complete and are less frequent.

Therapy Based Upon Evaluation of Factors Influencing Prognosis

Fact	Mild or Moderate	Sev	Very Severe
Age	Under 40	Over 40	
Organism count in sputum/oil field	Under 30	30-75	Over 75
Lob in oled	Singl	2 or 3	4 or 5
Platelet	Under 120	Over 120	Over 140
Blood pressure			
Syst li	Over 90	Under 90	Shock o
Diastol	Over 60	Under 60	pulmonary dema
Leukocyte count	Over 10,000	6,000-10,000	Under 6,000
Albumin	0 to ++	+++ to ++++	
Assessment	0 to mild	Moderate	Severe
Complications	Non or at al fusion	Empyem lung abscesses	Meningitis endocarditis
Blood culture	Negative	Positive	
Pneumococci type	High type	I II III IV VII	
Mortality rate			
Range	0-10%	25-10%	10-50%
Average	0-4%	4%	20%
Therapy indicated	Under dose of penicillin broad spectrum antibiotic or sulfonamides	High dose of penicillin broad spectrum antibiotic or sulfonamides	Max doses of penicillin

(Modified from Morris F. Collen, Penicillin Foundation Medical Bulletin VI 31 January 1948)

may be administered in several ways. The soft rubber facial mask of the BLB OEM of Bennett type is probably best. With this mask oxygen concentration up to 85% may be easily furnished. Oxygen tent is generally used if patient is intubated or tracheostomized who would otherwise remove the mask. However, the tent is generally advised because the average concentration of oxygen is only about 40-50% and unless watched carefully carbon dioxide may accumulate.

B. Fluid. Fluid intake must be decided whether given orally or parenterally to maintain a urine output of at least 1500 cc. Patients taking sulfonamides should have sufficient alkalinizing powder so that the urine is above pH 7 at all times. Potassium bicarbonate should be used in patients with aural or potential heart failure. Beware of excessive potassium toxicity.

C. Diet. During the severe acute phase patients usually have little desire for food. During this acute phase food intake is of little importance. Patients who develop complications and have long convalescence should be placed on high protein high vitamin high calorie diet.

Symptoms and Supportive Measures

A. Toxemia. The treatment and activity of the toxin in the blood stream which may occur in severe pneumonia must be controlled.

3 Sulfonamides

- a Sulfadiazine Sodium Injection U S P or Sulfamerazine Sodium Injection U S P 5.0 Gm (75 gr) or sulfadiazine and sulfamerazine 2.5 Gm (37½ gr) of each, in 500 cc (1 pt) M/6 sodium lactate I V as soon as diagnosis is made
 - b At the same time give 4.3 Gm (60.75 gr) sulfadiazine sulfamerazine mixture or sulfisoxazole (Gantrisin®) orally
 - c Continue with 3.4 Gm (45.60 gr) sulfadiazine sulfamerazine mixture or sulfisoxazole (Gantrisin®) every 8 hours together with 1.2 Gm (15.30 gr) sodium bicarbonate with each Gm of sulfonamide. If patient is unable to take oral medication he may be maintained on 5 Gm (75 gr) sodium sulfadiazine or sodium sulfamerazine or mixture of the two in 500 cc M/6 sodium lactate I V every 8 hours watching the electrolyte balance. If alkalois develops give the sodium sulfadiazine or sodium sulfamerazine in a line. Do NOT administer sulfonamides in glucose, blood, plasma or amino acid solutions.
- C. V ery Seve re Ca ses Patients with v ery s eve re pn umonias should receive massive penicillin therapy in an attempt to achieve a pneumococcal concentration of penicillin in infected areas as rapidly as possible. Combinations of penicillin and broad spectrum antibiotics or sulfonamides offer no advantage over penicillin alone.
- 1 Penicillin 1 million units of aqueous penicillin I M every 2 hours or continuous administration of penicillin solution of 10-12 million units daily by I V or I M drip until favorable clinical response occurs
 - 2 Chlorotetracycline oxytetracycline or tetracycline 0.5 Gm I V every 12 hours or chloramphenicol or erythromycin 0.5 Gm every 6 hours until favorable response occurs
 - 3 Sulfonamides As for severe form Give 5.0 Gm (75 gr) sodium sulfadiazine or sodium sulfamerazine or of a mixture of the two I V at once and follow with oral or I V maintenance as indicated. Always maintain adequate alkalization and fluid intake

Evaluation of Therapy

If there is no response to therapy in 24 to 36 hours completely re-examine cases for cause

- A Incorrect Diagnosis Infection may be caused by microorganism resistant to anti microbial agent used. Where etiology of pneumonia is in doubt broad spectrum antibiotics are usually preferable to penicillin
- B Spread of Pneumonia Treat as for erysipeloid. If not already doing so substitute one of the tetracyclines or chloramphenicol for therapy already under way
- C Development of Complications 1 empyema lung abscess endocarditis meningitis
- D Presence of an associated condition that may give fever

General Measures

- A Oxygen is very important in any patient with severe or moderately severe pneumonia cyanosis or marked hypoxia and

from the intestines

- 2 Neostigmin Methylsulphate U.S.P. (1:2000 Sol.) 1 cc (15 mg) subcut and insertion of a nasal tube will generally produce rapid initial decompression
- 3 Stomach tube for dilatation of the stomach Suction through a nasal tube passed into the stomach is unnecessary

Complications

- 1 Congestive failure In elderly patients or patients with preexisting heart disease congestive failure may be precipitated by the pneumonia. When this occurs digitalization by one of the rapid methods is indicated (see page 197). This must be distinguished from shock and pulmonary edema (see page 126).
- 2 Cardiac arrhythmias The occurrence of extrasystoles is common and generally requires no treatment. If auricular fibrillation, flutter develops rapid failure may be precipitated. Digitalization by one of the rapid methods is generally indicated in these cases (see page 197).

Complications

For treatment of these complications see the respective diseases (Modified after Collen)

Complication	% Incidence
Stippled pleural effusion	4.5
Empyema	0.3
Lung abscess	0.3
Pneuritis	0.3
Endocarditis	0.1
Meningitis	0.1

All pleural effusions associated with pneumonia must be aspirated promptly if they are empyemic which may be indicated (see page 141).

STREPTOCOCCIC PNEUMONIA

(Lobar code No 360 102) (Bronchopneumonia code No 361 102)

An uncommon type of pneumonia, usually secondary to a preceding pulmonary infection (i.e. viral pneumonia influenza, etc.). Onset is most often gradual but is at times sudden with severe intoxication marked dyspnea and cough with bloody or mucopurulent sputum. Pleural effusion occurs early and is fairly common and may progress to empyema. Most cases are due to *Streptococcus pneumoniae*.

Physical findings vary with severity. There may be only slight dullness and moist rales. In severe cases pleural effusion obscures pneumonia signs. Thorax is usually reddened and has some xeroderma.

Treatment

Penicillin is the drug of choice. Dose is similar to that for pneumococcal pneumonia.

to save the patient from exhaustion or circulatory failure

1 Paraldehyde is the drug of choice for this purpose

- Oral 8 cc (2 dr) at once if there is no response in 30 minutes repeat dose until patient is quiet Then give 4 15 cc (1 4 dr) every 3 4 hours p r n restlessness
- Intramuscular If patient is unable to swallow give 4 cc (1 dr) at once if there is no response in 30 minutes repeat dose until patient is quiet then give 4 8 cc (1 dr) i M every 3 4 hours p r n restlessness

2 Barbiturates Mild restlessness and sleeplessness can be treated with the following drugs

- Phenobarbital sodium (phenobarbital sodium) 0 1 Gm (1 1/2 gr) at bedtime
- Phenobarbital (phenobarbital) 15 30 mg (1/4 1/2 gr) t i d during the day

B Shock and Pulmonary Edema The usual cause of death in pneumonia is shock (circulatory collapse) and/or pulmonary edema The most important factor in management of these conditions is very early recognition and prompt vigorous treatment Therapy is essentially the same in both conditions

1 Treatment of SHOCK see page 27

2 Treatment of anoxia Because of the important role of anoxia in the production of shock and pulmonary edema in pneumonia prompt initiation of oxygen therapy preferably with positive pressure facial mask is of utmost importance

C Cough Generally requires little therapy and clears spontaneously with adequate treatment Expectorants are of little value if cough is severe Codeine may give relief and help patient to sleep

1 Codeine phosphate 0 03 0 06 Gm (1/2 1 gr) orally or subcutaneously every 3 4 hours

2 R Ephedrine sulfate

or hydrochloride (1% Sol) 30 0 31

Codeine phosphate 0 5 gr vials

Syrup of wild cherry or

other vehicle q d 120 0 3 v

Sig 4 (1 t p) q 3 4 hours p r n cough

D Pleuritic Pain

1 Mild pain Ethyl chloride spray for cutaneous anesthesia Spray for about 1 minute over area of pain and vertically over entire area of pain so that a line of frosting is visible The method gives relief in pain for 1 to 10 hours in greater majority of patients

2 Severe pain Procaine hydrochloride infiltration (1% 1% Sol) is injected subcut in a vertical line passing through the area of greatest pain and 5 cm higher and lower

3 Very severe pain Meperidine Hydrochloride U S P (Pethidine Hydrochloride B P (Demerol®) 50 100 mg appears to be drug of choice for treatment of pleuritic effect on the respiratory and cough reflexes

E Abdominal Distention Abdominal distention generally disappears with swallowing associated with effective pneumonia is frequent in patients with pneumonia

1 Oxygen therapy in high concentration (90 100%) is usually most useful because the oxygen is very rapidly absorbed

from the intestine

- 2 Nitroglycerine Methyl sulfate U S P (12000 Sol) 1 cc (15 mg) about 1/2 inch of a rectal tube will generally produce spontaneous decompression
- 3 Stomach tube for dilatation of the stomach. Suction through a nasil tube passed into the stomach is necessary

Fluid Abnormalities

- 1 Congestive failure. In elderly patients or patients with pre-existing heart disease congestive failure may be precipitated by the pneumonia. When this occurs digitalization by one of the old method is indicated (see page 197). This must be distinguished from shock and pulmonary edema (see page 186)
- 2 Cardiac arrhythmias. The occurrence of extrasystoles is common and generally requires no treatment. If auricular fibrillation or flutter develops rapid failure may be precipitated. Digitalization by one of the older methods is generally indicated in these cases (see page 197)

Complications

For treatment of the complications see the respective diseases (Modified after Collin)

<u>Complication</u>	<u>% Incidence</u>
Striopulural effusion	4.5
Empyema	0.3
Lung abscess	0.3
Pericarditis	0.3
Endocarditis	0.1
Meningitis	0.1

All pleural effusions associated with pneumonias must be tapped promptly to detect early empyema which may be treated medically (see page 141)

STREPTOCOCCIC PNEUMONIA

(Lobar code No 360 102) (Bronchopneumonia code No 361 102)

An uncommon type of pneumonia usually secondary to a preceding pulmonary infection (i.e. virus pneumonia, influenza, measles). Onset is most often gradual but is sometimes sudden with severe intoxication marked dyspnea and cough with bloody or mucopurulent sputum. Pleural effusion occurs early in fairly common and may progress to empyema. Most cases are due to patholytic streptococci.

Physical findings vary with severity there may be only at red dullness and moist rales. In severe cases a pleural effusion obscures pulmonary signs. Thorax is usually reddened and has a mucous exudate.

Treatment

Penicillin is the drug of choice. Dosage is similar to that for pneumococcal pneumonia.

STAPHYLOCOCCAL PNEUMONIA (code No 381 105)

An uncommon type of pneumonia usually secondary to preceding infection. The onset is usually gradual and progressive to grave illness. Cough and dyspnea are common. Multiple lung abscesses occur frequently. Patchy consolidation with diffuse rales are often found. Sputum is variable in appearance.

Treatment

Sensitivity tests should be performed. Pending the results of the tests the following should be given: *I.M.* every 6 hours Erythromycin U.S.P. (Erythrocin®) 0.5 Gm. Novobiocin N.N.D. (Cathomylin® Albamycin®) 0.5 Gm. Chloramphenicol U.S.P. (Chloromycetin®) 0.5 Gm. Bacitracin U.S.P. 20 000 units.

FRIEDLANDER'S PNEUMONIA (code No 381 131)

Pneumonia due to *Klebsiella pneumoniae* is often associated with chronic debilitating diseases. The onset is usually sudden with chills, fever, dyspnea, cyanosis, cough and marked toxicity and in most cases progresses rapidly to a fatal termination. There is a tendency to necrosis and abscess formation in the subacute or chronic forms. Early recognition is imperative for favorable outcome.

Physical findings are variable and extensive involvement may give only dullness and diminished breath sounds. Sputum is reddish mucoid and tenacious giving a currant jelly appearance. White blood cell count is variable; may have leukopenia or leukocytosis.

Treatment

A Specific Measures Treatment of Friedlander's pneumonia is very severe infections.

1 Streptomycin 1 Gm. every 6 hours until favorable response occurs then 0.5 Gm. every 6 hours until afebrile 3 days and 2 One of the tetracyclines (*I.M.* or *I.V.*) or chloramphenicol 0.5 Gm. every 6 hours or Gentamicin® 1 Gm. every 6 hours. Continue for 2 weeks.

B General Measures See Pneumococcal Pneumonia page 124.

HEMOPHILUS INFLUENZAE PNEUMONIA (code No 381 110)

A rare form of pneumonia which usually is rapid in onset and progression. The outstanding features are severe inflammation of the bronchi and bronchioles leading to bronchiectasis and hemorrhagic edema of lungs. Patients are extremely toxic.

There is a patchy consolidation and the sputum is bloody. Leukopenia is frequently present.

Treatment

A Specific Measures Continue treatment for 10 days after temperature has returned to normal.

1 Combined streptomycin and sulfonamide therapy is the treatment of choice. Streptomycin 0.5-1 Gm. every 6 hours *I.M.*

and if named as follows: v r y s v e p n m o a l p e u
mon (page 124)

2 Combination of cylin s (I M o I V) o chl mphe ol
0.5 Gm v y 6 hou s plu s l f onamid as fo ever
p mo cal p um n i s

B Ge e l M S P e moc i P mo ia p ge 124

PRIMARY ATYPICAL (VIRUS) PNEUMONIA (code No 360 160)

Virus pneumonia is most commonly has a gradual onset with chill
n s mild fever non productive cough malaise and headache
The patient usually does not appear too ill and for this reason the
disease in most cases is not recognized

The physical findings are variable may be absent or how
small areas of r pitant ral and occasional dullness The fever
reaches peak in 2-3 days and falls by lysis for the next 10 days
In every case fever may last for weeks

Röntgen examination of chest reveals patchy consolidation
The laboratory findings are usually within normal limits Cold
agglutination test positive in about 50% of cases throughout
the

Treatment

A Supportive Measures Give oxygen if the following lines 0.25-0.5

Gm every 8 hours orally I V the apy may be necessary in
every case or if the patient is vomiting may be combined with
oral therapy as tolerated Give 0.5 Gm ery 12 hours

B Ge e l M S P e moc occ Pne mon a p g 124

PULMONARY TUBERCULOSIS (code No 360 123)

Diagnosis

One of the principal problems in the diagnosis of tuberculosis
the finding of early cases before symptoms are present The only
way to do this early case is by periodic x-ray examinations of the
chest When the characteristic symptoms of cough weight loss
and night sweats appear the diagnosis is usually made
certain step that must be followed in making a diagnosis of a tubercu-
lar pulmonary tuberculosis

A Diagnosis of a Pulmonary Lesion Always check positive
x-ray film for any possible New assay accepted
he does not have pulmonary tuberculosis without a positive
examination of the chest

B Proof of Tuberculosis

1 Skin test The intradermal method is the most reliable B
gun with 1:10,000 O.T. if this is negative give 1:1000 if
this is negative use 1:100 In single PPD test with last
test if this is negative use 2-d t gth The test is
read in 48-72 hours and at least 5 mm induration (with or
without redness) is required for a positive test

With few exceptions a negative reaction to tuberculin in the dilution of 1:100,000 or weaker strength does not exclude tuberculosis

- 2 Recovery of tubercle bacilli This is the only certain method of establishing the diagnosis. However, this may occasionally be difficult and it may be necessary to institute treatment on the basis of a strong presumptive diagnosis (i.e., a typical x-ray lesion in a young person with a positive skin test who has been in contact with an active case). In the absence of sputum, a gastric or bronchial lavage may be necessary. Bronchial lavage is done by spraying the pharynx and hypopharynx several times with 1% Pontocaine®. The patient should be fasting in order to avoid aspiration if emesis occurs. With the aid of a laryngeal mirror, 5-10 cc. normal saline is injected between the vocal cords via a curved cannula. The patient coughs this out into a sterile container. Injection of saline may have to be repeated several times to get an adequate specimen (approximately 15 cc.). The specimen may be examined for tumor cells as well as for tubercle bacilli. The patient must not take anything by mouth for two hours after the procedure.

Non-pathogenic acid fast bacilli may occasionally cause confusion, especially in smears of gastric contents. Guinea pig inoculation will differentiate these from tubercle bacilli. When tubercle bacilli cannot be recovered, other diagnostic possibilities must be considered.

- C Activity of a Tuberculous Lesion Once tuberculosis has been diagnosed, the status of the lesion must be determined to decide if therapy is needed.

- 1 Presence of tubercle bacilli in the sputum is evidence of activity.

Either progression or regression of the lesion by x-ray especially within short periods of time indicates activity.

- 3 A cavity is usually evidence of activity.

- 4 Other manifestations suggesting activity

- | | |
|-----------------------------------|--------------------------------------|
| a Afternoon fever | e Pleurisy with or without effusion |
| b Night sweats | f Increased blood sedimentation rate |
| c Blood streaked sputum | |
| d Weight loss and/or fatigability | |

- 5 Any newly discovered tuberculous lesion should be considered possibly active. If this cannot be immediately determined, sputum studies should be continued together with close observation by monthly x-rays for 2-3 months. If no change occurs and ultas for tubercle bacilli remain negative, the interval between films can be lengthened, but observation should be continued for at least 2 years.

Treatment

- A Rest Bed rest and mental relaxation in cheerful surroundings, either at home or in a sanatorium, should be instituted whenever an active lesion exists or is probable. This is still an important measure in the therapy of pulmonary tuberculosis. Although the duration of the rest period required has been reduced by the antituberculosis drugs, bathroom privileges may be permitted where symptoms are minimal or absent.

- 1 Advantages of sanatorium care. Most authorities prefer sanatorium care to home treatment. The several advantages to institutional management:
 - a Well controlled and symptomatic environment while the chief occupation is the care of tuberculous patients.
 - b Educational instruction in the care of tuberculosis by means of a well organized program.
 - c Contact with other persons who have the same disease and in the problems. Observation of the success of a well controlled program of rest.
 - d Complete break with old environment and more complete mental and physical rest.
 - e Medical observation more frequent and control of management more complete.
 - f Planned program of gradual increase of activity and recreation under well trained personnel.
 - g Active therapy (effective drug, collapse, surgical, etc.) more easily administered.
 - h Isolation program can more easily be enforced thus preventing the spread of the disease.

- 2 Advantages of home care. Home care is sometimes desirable but safety of other family members undisturbed rest period administration of medicine etc. must be provided for. Advantages and indications:
 - a Local facilities lead to a private patient.
 - b For patient unable to adjust to institutional care.
 - c Attended home treatment can sometimes be stabilized by the mere presence of the patient.

B. Drug Therapy. This has become the most important measure in tuberculosis treatment. It is indicated in all cases of active disease and is most effective when administered in conjunction with a well regulated program of bed rest as well as collapse therapy when indicated (see below). In general the return to limited physical activity is permitted sooner with drug therapy and gradual ambulation may be started as soon as clinical improvement is well established.

Duration of drug therapy in pulmonary tuberculosis. The pre-nitro recommendation is for prolonged administration of combination of the drugs isoniazid (INH) and streptomycin (a dipeptide amide). Many patients seem to be efficient for prolonged treatment even after moderate resistance of the organism to the drugs has been shown by sensitivity tests. Most authorities advise a minimum of 12 months of drug treatment after the final statistics (National Tuberculosis Association 1955) has been attained. It should be emphasized again that drug treatment has been shown to be a substitute for rest and collapse therapy.

The principal drugs now used in the treatment of pulmonary tuberculosis are isoniazid (INH), streptomycin or dihydrostreptomycin and amino glycolic acid (PAS). The simultaneous use of these two drugs is probably justifiable for severely ill patients but in the more chronic forms of pulmonary tuberculosis no definite advantage has been shown. In general it is probably wise to withhold isoniazid or streptomycin preferably the latter if possible later in the course of treatment.

- 1 Isoniazid U S P (INH) (See page 133 for dosage) This is the most effective drug currently available. However when used alone its effectiveness is decreased by the early development of bacterial resistance. It should be used with at least one of the other drugs mentioned below.
 - a Indications Any active tuberculous lesion including primary tuberculosis in children. This drug has particular value in military tuberculosis, tuberculous meningitis (see page 468), streptomycin resistant tuberculosis and streptomycin intolerance.
 - b Toxicity Toxic reactions to isoniazid are infrequent in the usual dose of 5 mg/kg/day. They include dermatitis and fibrotic reactions. With larger doses peripheral neuropathy and rarely CNS irritability may occur. There is good evidence that the latter are related to pyridoxine depletion. Supplementary doses of pyridoxine (25-50 mg/day) should be given.

2 Streptomycin Sulfate U S P and Dihydrostreptomycin Sulfate U S P (See page 133 for dosage)

- a Indication The indications for these drugs are the same as for isoniazid except that they are less effective than isoniazid in advanced tuberculosis. Like isoniazid they are less effective when used alone and whenever possible should be given in combination with at least one of the other drugs.

- b Toxicity Streptomycin and dihydrostreptomycin are essentially like in the therapeutic effect. Since the toxicity of dihydrostreptomycin for the eighth nerve (deafness) is more marked than that of streptomycin (vertigo) the latter is more commonly used. Experience has shown that dividing the usual dose into equal parts of streptomycin and dihydrostreptomycin reduces the toxicity of both drugs.

Toxic reactions to these drugs are few when given twice weekly. There is ample evidence that the regimen produces a therapeutic effect comparable to other streptomycin schedules (except in the more serious forms of the disease where daily dosage may be necessary). Generalized dermatitis occasionally occurs in children and the dosage must be stopped. Perioral numbness often appears shortly after injection and may last for several hours. By itself it can be ignored.

3 Aminosalicylic Acid U S P (para-aminosalicylic acid PAS) (See page 133 for dosage)

- a Indications This drug has a low level of antibacterial activity but when used with streptomycin or isoniazid it delays the emergence of resistance.
- b Toxicity Toxic reaction to PAS includes nausea, vomiting and diarrhea a few days after the start of therapy. Generalized dermatitis. The gastrointestinal symptoms may sometimes be overcome by gradual increase in dosage or by topping the dosage for a few days and then bringing it in small doses gradually increasing to the regular dose in 2-3 weeks. When necessary dermatitis due to PAS occurs the dosage usually must be stopped.

4 Viomycin Sulfate N N D (Viactane® Vioin®) and

effective and more toxic drug has limited usefulness when hemotherapy is indicated and the above mentioned drugs cannot be used. The usual dose is 2 Gm 1 M daily or twice weekly for up to 6 weeks.

5. Pyriminamide (pyrazinamide PZA) a drug which occurs as a lipoid soluble toxic hepatitis is not used alone with isoniazid for periods of 13 months when there is a hyper sensitivity to the other drugs present. Clinical observations for symptoms and laboratory evidence of liver dysfunction must be ruled out and the drug stopped promptly if any abnormality appears. The usual adult dose is 0.75 Gm twice daily.
6. Cycloserine (N N D (S omic®) The place of this drug in the treatment of tuberculosis has not yet been determined.

ANTI TUBERCULOSIS DRUGS

DRUG	ADULT DOSE		REMARKS
Streptomycin Dihydrostreptomycin	1 Gm daily or 1 Gm 2 times per week		Only indicated for treating these drugs usually is hypersensi- tivity of the patient or known resistance of bacilli to other drugs
Aminosalicylic Acid (PAS)†	4-5 Gm tid po		
Isoniazid (INH)	5-10 mg per Kg per day		
Combined Therapy	Cotrimoxazole	Intermittent	Any two of these the drug may be used (except in very severe cases all three are indi- cated) U INH
Streptomycin	1 Gm per day	1 Gm twice weekly	where possible Isoniazid is daily treatment in (till improvement is established then twice weekly) and 10 mg per Kg per day of INH
Aminosalicylic Acid	4 Gm tid po (with 1/3 of above schedule)		
Isoniazid	5-10 mg per Kg per day (with the above schedule)		

Equal part of streptomycin dihydrostreptomycin (e.g. 0.5 Gm of each) as a substitute with the other drugs.

†Pyriminylpyrazinamide aminosalicylic acid Mycobacterium or sodium salicylic acid.

C Collapse Therapy Collapse therapy is often of benefit in the treatment of recent lesions (especially where cavitation is present) permitting earlier ambulation. Pneumoperitoneum is the principle method now used, having largely replaced pneumothorax because of the high incidence of irreversible pleural complications in the latter. Once instituted treatment should continue for 3-4 years for maximum effectiveness.

D Surgery

1. **Pulmonary resection** This has gained increasing popularity in the treatment of pulmonary tuberculosis in recent years although only about 10-15% of patients now being treated in tuberculosis hospitals require major surgery. The following indications are widely accepted:

- Localized nodule especially where diagnosis is in doubt
- Bronchiectasis causing positive sputum
- Bronchostenosis
- Thoracoplasty failure (Some of these can be successfully treated by resection)
- Any localized chronic focus which has not become inactive (National Tuberculosis Association 1955) after 9-12 months of adequate nonsurgical therapy

2. **Thoracoplasty** The current indications are decreasing in number and are as follows:

- Chronic cavity lesions where resection is not feasible and where the lesser procedure can be tolerated
- Certain cases where later resection is contemplated and it is felt that thoracoplasty will improve the patient's general condition
- To reduce pleural dead space after a large pulmonary resection and thus minimize overdistention of remaining lung tissue
- To close chronic empyema spaces

E Diet A diet adequate in calories and high in proteins and vitamins should be employed. One should generally attempt to keep the tuberculous patient's weight above normal. No specific diets have been shown to be of benefit. If the patient is not eating and is losing weight, forced feedings by tube if necessary should be instituted.

F Climate There is little evidence that climate plays an important part in the management of tuberculosis. In the past heliotherapy or ultraviolet irradiation was widely recommended. There is no evidence that this therapy is of any value and it may be harmful in pulmonary tuberculosis. A mild sunburn or direct sunlight to the chest.

G Symptomatic Treatment Patient should be reassured that his symptoms will disappear as the illness is brought under control.

1. **Cough** In general cough in tuberculosis should not be suppressed by means of drugs. The nonproductive cough serves no useful purpose and can generally be controlled voluntarily. If cough is productive the patient should be encouraged and instructed to cough properly (i.e. without initial violent inspiratory phase; the actual cough likewise should be without effort). It may become necessary at times to use medication to suppress a cough when it is exhausting to the patient. In these cases codeine 8-15 mg ($\frac{1}{8}$ to $\frac{1}{4}$

g) or lly may be helpful. Bonyai has found that int mit
t tinal tio f 5-10% carb n dioxide with o ygen ca ses
a d minution of the co gh. Ca es w th la ge cavities and
copious sput m may b helped by po tural d sinage.

- 2 Night sweat Effo ts t cont lth should be d cted at
a oid g exc iv nd unnec ry bed clothing. Atr pi
e lf te 0.4-0.6 mg ($\frac{1}{150}$ - $\frac{1}{100}$ gr) ally may be t led
at b d tim but i g n ally of l tle val e.

- 3 Hemorrhage The chief da ge f om h morrhage tub
culos is not s dden de th b t asp rati of the inf ted
bl od a d ap e d of th disea e to other p rts f the lung.
Th efo e do not u e c gh inhibit s in the tre tm nt of
hem rhage *morphine sulfate should be avoided*.

Sh k the apy hould b i stit ted if bleed g i sev e
and shock is imm n nt (see pag 27).

- b R ass an is most import ti allaying appr he sio

c Phenob rbtal sod um 60-120 mg (1-2 g) s bc t may
be of val e in q iet g th appr hens e patient.

- d Collap the apy. At tim if s vere bl ding onti ues
collap e th py may b com an m rgen ym a ur b t
th s in l es the da ger of cau ing pr ad of the disea e.
Whe e e bleed g co t ues. Poterio Pituitary In
j tio U S P 1 cc (10 I U) in 10 cc orm l salin
can be given SLOWLY I V (1 cc /min). Bleedings me
tim stop promptly.

- f N nsp if c mea res. Ab ol t bed rest is e ential.
The v l e of position g is ont ov sial but c mplet
mmobilizati n unwi e. Mo ing f om tim to tim
h lp b i g up e et ns. Instr t patient in pr per
m thod of c ghi g (ee abov).

P ophyla is

A l l tl P s tio. All nece ryp eca tio m t be taken
wh n se i found o s p e ted.

- B BCG Imm l at. Although ext s v st des e till i
p ogr the val e of cti e mmuni tion of the ge e al pop
ulati n with BCG v ci e r m i do btful afte many y a s.
U til the r lt a e va labl ro ti immunizati ns hold
not be considered exc pt fo ind vid al w th gat ve t b c l n
t ts in wh m the ri k of i f tio is g eat (i e m d l
st dent n es etc).

- C T atment of R e t T b e c lin Co rto. Mo t a thor ties
ow ec mm d the t e tment of c r t in ind vid al kn wn to
have b en e tly (withn the p ced g 6 month) infected by
t b lo is ven th gh a e iden e of the disea oth th a
p s itiv tub r ulin rea t o is p ent. Child en up t ag 3
should routi ly re eiv a o r e of d g t eatme t (s e b low)
when v a positiv tub ulin e ti is fo d i c it is in
this group that the mo t ri s mplicatio of tub r losis
e mo t likely to oc ur. Some ad o at th tr atm t of
ce t t b culin on rt re f a y age.

Thi p e ti tr atm nt o sist of a u of dr g
th apy with t oth r r t i tio. Isoni rid 5 mg /Kg /d y
should b d tog ther with min ali yli a id (PAS) 200
mg /Kg /day f 8-12 mo ths. Whe e PAS is n t tol t d
st ept my in 20 mg /Kg twi e weekly m y be batit t d.

LUNG ABSCESS (code No 360 100 2)

The etiology of lung abscess should be determined in all cases before therapy is instituted because the underlying process may be as important as the abscess and may modify the course and treatment. About 25% of the cases follow oropharyngeal surgery. Causes of the remainder include pneumonia, aspiration of foreign body, bronchogenic carcinoma or other bronchial obstruction, bronchiectasis, and emboli.

Diagnosis

A History

- 1 Acute lung abscess Usually follows oropharyngeal surgery, pneumonia, or other systemic or local infections. There may be a latent period, especially post surgical, followed by rapid development of cough, malaise, chills, and fever, and pleurisy with or without effusion. Patient becomes very toxic and breath becomes foul. Several days later patient may suddenly cough up a large amount of pus, and at this stage the diagnosis is certain.
- 2 Chronic abscess The manifestations vary with extent and location of the abscess. There is generally low grade fever, cough with copious sputum, pleurisy, and weight loss. Chronic lung abscess usually occurs secondary to bronchiectasis or bronchial obstruction.

B Physical Examination Findings vary with the size, position, contents of the cavity, and local pulmonary reaction.

C Laboratory Findings

- 1 Histological examination usually typical of severe infection.
- 2 Chest x ray may show cavitation with a fluid level and surrounding pneumonitis.

D Bronchoscopy This is an important diagnostic procedure inasmuch as an obstructive lesion or foreign body in the bronchus may be found.

Treatment

A Specific Measures

- 1 Acute abscess Begin antibacterial agents early to prevent destruction of lung tissue. Make smear and culture of sputum to determine predominant organism, and employ the appropriate antibacterial agent(s) in very high doses (see table on page 514). If patient fails to respond, surgery is indicated without delay.
- 2 Chronic abscess Although some few chronic cases will get well on a medical regimen as outlined above for acute abscess, therapy with antibiotic agents is used to decrease infection in preparation for surgery.

B General Measures

- 1 Supportive and symptomatic care
- 2 Postural drainage is very important to prevent accumulation of material (see page 121).
- 3 Do not depress the cough reflex by use of sedative cough medicines.
- 4 Bronchoscopy is usually indicated to promote drainage.

C Follow up The patient cannot be considered cured until there

has been completed imaging of the lung by x-ray. Serial x-rays required in this follow-up which may take many weeks.

NEOPLASMS OF THE LUNGS BRONCHOGENIC CARCINOMA (code No 350.8) (Epidermoid code No 350.814)

Neoplasms of the lung form a very important group of the malignancies. The most static tumors are the most common but the primary neoplasms are of great interest in diagnosis and therapy. The primary tumors usually arise from the bronchus and spread into the lung field. They are rarely diagnosed early because of the insidious onset and tendency to mimic other pulmonary diseases. Characteristically the common presenting symptom is an asthmatic process, progressive cough becomes productive and hemoptysis on consolidation. Atelectasis, lung abscess, and pleural effusion may occur. It must always be considered in the diagnosis of any acute or chronic pulmonary disease especially in males over 50 years of age. Bronchoscopy and examination of sputum for cancer cells are important diagnostic studies.

Treatment

- A. Surgery: the treatment of choice when the lesion is discovered early.
- B. Supportive and symptomatic measures if the cases in which surgery cannot be performed.

PULMONARY ATELECTASIS (Compression code No 362.435) (Postoperative code No 362.415.4)

Atelectasis is a partial or complete obstruction of the bronchus with subsequent collapse of the lung distal to the obstruction. Most cases follow major surgery and tend to occur in the right lower lobe. The condition usually is manifested 4 days after surgery and the finding is the collapse of the entire lobe and collapse of the involved segment. If immediate treatment is not carried out, secondary bacterial pneumonia and a pneumonia develop.

Treatment

- A. Postoperative Atelectasis
 1. For prevention of hypoxia, the volume of oxygen by use of a mixture of 93% O_2 and 5% CO_2 administered by mask for several minutes every 1-3 hours. This is also good preoperative measure.
 2. Bronchodilation by low intermittent positive pressure (e.g., Bennett valve) has been demonstrated to be most effective. The pressure should be reduced to 30 mm Hg every 2-3 hours for 24 hours before discharge that the mechanical cause is removed.
 3. Aspiration of the collapsed lobe with the use of the aspirator placed blindly through the nasopharynx with the aid of a laryngoscope if effective.

LUNG ABSCESS (code No 380 100 2)

The etiology of lung abscess should be determined in all cases before therapy is instituted because the underlying process may be as important as the abscess and may modify the course and treatment. About 25% of the cases follow oropharyngeal surgery. Causes of the remainder include pneumonia, aspiration of foreign body, bronchogenic carcinoma or other bronchial obstruction, bronchiectasis and emboli.

Diagnosis

A History

- 1 Acute lung abscess Usually follows oropharyngeal surgery, pneumonia or other systemic or local infections. There may be a latent period, especially post surgical, followed by rapid development of cough, malaise, chills and fever and pleurisy with or without effusion. Patient becomes very toxic and breath becomes foul. Several days later patient may suddenly cough up a large amount of pus and at this stage the diagnosis is certain.
- 2 Chronic abscess The manifestations vary with extent and location of the abscess. There is generally low grade fever, cough with copious sputum, pleurisy and weight loss. Chronic lung abscess usually occurs secondary to bronchiectasis or bronchial obstruction.

B Physical Examination Findings vary with the size, position, contents of the cavity and local pulmonary reaction.

C Laboratory Findings

- 1 Histological examination usually typical of a pyogenic infection.
- 2 Chest x ray may show cavitation with a fluid level and surrounding pneumonitis.

D Bronchoscopy This is an important diagnostic procedure inasmuch as an obstructive lesion or foreign body in the bronchus may be found.

Treatment

A Specific Measures

- 1 Acute abscess Begin antibacterial agents early to prevent destruction of lung tissue. Make smear and culture of sputum to determine predominant organism and employ the appropriate antibacterial agent or agents in very high doses (see table on page 514). If patient fails to respond surgically indicated without delay.
- 2 Chronic cases Although some few chronic cases will get well on a medical regimen as outlined above for acute abscess, therapy with antibiotic agent is used to decrease infection in preparation for surgery.

B General Measures

- 1 Supportive and symptomatic care
- 2 Postural drainage is very important to prevent accumulation of material (see page 121).
- 3 Do not depress the cough reflex by use of sedative cough medicine.
- 4 Bronchoscopy is usually indicated to promote drainage.

C Follow up The patient cannot be considered cured until the

prophylaxis. Although it is occasionally beneficial, it has the disadvantage of increasing total alkalosis. The usual dose is 250 mg (4 g) twice daily.

PULMONARY INFARCTION

(Due to Thrombus code No 360 511)
(Due to Embolus code No 360 512)

Pulmonary infarction due to an embolus thrombus occluding a branch of the pulmonary artery. A wedge-shaped area of lung tissue distal to the occlusion becomes infiltrated with blood which extravasates into cells and pulmonary tissue.

Diagnosis

The signs and symptoms vary with the size of the blood vessel occluded. If the infarct is small there may be no manifestations. If large the most common findings are hemoptysis, chest pain (usually pleuritic) and dyspnea. If massive enough there may be right heart failure and death. If patient survives moderate or severe infarction dyspnea, chest pain, pleurisy with or without effusion and fever become prominent. ECG evidence of right heart strain may be a very early finding. Leukocytosis and increased sedimentation rate persist until the infarction has healed. X-ray of the chest may show wedge-shaped areas of consolidation.

Treatment

When a patient has a pulmonary embolus suspected without thrombosis and institute immediate therapy for the thrombosis (see page 215).

A Emergency Measures

- 1 Oxygen in high concentration preferably 100% (by mask) to overcome asphyxia. This also helps prevent cardiorespiratory failure.
- 2 Morphine 30-60 mg ($\frac{1}{2}$ to 1 gr) and atropine 1-2 mg ($\frac{1}{60}$ to $\frac{1}{30}$ gr) I.V. slowly and repeat every 3-4 hours. This helps overcome the general pulmonary arteriolar spasm that occurs.
- 3 Morphine sulfate 8-15 mg ($\frac{1}{8}$ to $\frac{1}{4}$ g) or Meperidine Hydrochloride U.S.P. (Demerol®) 50-100 mg ($\frac{3}{4}$ to $1\frac{1}{2}$ g) subcutaneous or I.V. to help control pain.
- 4 If shock persists I.V. fluid should be used with great caution because of possibility of precipitating acute right heart failure.
- 5 If acute right heart failure develops venesection should be performed (see page 185).

B Follow-up Treatment

- 1 Observe carefully for evidence of infection and institute antibiotic treatment promptly if suggested.
- 2 Thoracic tests. If pleural effusion occurs and embolism respiration moves by physical signs.

Prophylaxis

The prophylaxis of pulmonary infarction consists of the treatment of the venous thrombosis (see page 217).

138 Pulmonary Emphysema

- 4 If the above fail aspiration of mucus by bronchoscopy is indicated
- 5 Antibiotic therapy Penicilline penicillin complex 300 000 units b.i.d.

B Spontaneous Atelectasis Bronchoscopy to determine the nature of the obstruction and then institute appropriate treatment

PULMONARY EMPHYSEMA

(Due to Unknown Cause code No 362 9x6)

(Postural code No 362-434)

Pulmonary emphysema is a disease usually found in older individuals and those suffering from chronic bronchial asthma. There is progressive distention of the pulmonary alveoli with subsequent rupture of the interalveolar membranes and replacement of the alveoli by larger poorly functioning air sacs.

The diagnosis is generally not difficult. The chief complaint is dyspnea. The anterior-posterior diameter of the chest is generally enlarged. The lung fields are hyperresonant and breath sounds coarse. Pulmonary emphysema must be differentiated from dyspnea due to congestive failure.

Treatment

A Specific Measures Since many patients have an associated chronic bronchitis with some elements of asthma therapy is generally similar to that outlined for chronic bronchitis or chronic bronchial asthma (see page 116)

- 1 Spasmolytic agents to relieve bronchial spasm. Epinephrin by inhalation, ephedrine etc (see page 118)
- 2 Eradicate any infection. Penicillin or tetracycline aerosols (see page 153)

B General Measures

- 1 Inhalation of 100% oxygen for 20-30 minutes helps relieve dyspnea and appears to reduce spasm. Recently the use of intermittent positive pressure in some cases with bronchial dilators has been shown to be of considerable value. Oxygen must be used with caution in cases with long-standing anoxia since use of 100% oxygen has caused coma and even death in some patients (see page 144)
- 2 Maintain mechanical efficiency of diaphragm at its optimum
 - a Abdominal belt. Obese patients or those with a poor abdominal musculature should wear an abdominal belt during the day (e.g. Kerr-Lag Noble)
 - b Manual relief of overdistention of lungs. The palms of both hands are placed under the anterior ribs and pushed inwards and upwards during end of expiration. This is repeated 10-15 times 2-3 times daily. Patients often claim their dyspnea is relieved for hours by this maneuver.
 - c Therapeutic pleuroperitoneum can be employed in selected patients to mobilize the diaphragm.
- 3 Acetazolamide (N.N.D. (Diamox®)) a bicarbonate dihydrase inhibitor has been used to reduce the bicarbonate content and the arterial pCO_2 in patients with CO_2 retention and

respiratory acidosis. Although it is occasionally beneficial, the disadvantage of increasing total acidosis. The usual dose is 250 mg (4 gr) twice daily.

PULMONARY INFARCTION

(Due to Thrombus code No 360 511)

(Due to Embolus code No 360 512)

Pulmonary infarction is due to an embolus or thrombus occluding a branch of the pulmonary artery. A wedge-shaped area of lung tissue distal to the occlusion becomes infarcted with blood which extravasates into cells and pulmonary tissue.

Diagnosis

The signs and symptoms vary with the size of the blood vessel occluded. If the infarct is small there may be no manifestations. If large the most common findings are hemoptysis, chest pain (usually pleuritic), and dyspnea. If more severe there may be right heart failure and death. If patient survives moderate or severe infarction dyspnea, chest pain, pleurisy with or without effusion, and fever become prominent. Example: evidence of right heart strain may be a very early finding. Leukocytosis and increased sedimentation rate persist until the infarct has healed. X-ray of the chest may show a wedge-shaped area of consolidation.

Treatment

When a patient has a pulmonary embolus suspect venous thrombosis and institute immediate therapy for thrombosis (see page 215).

A Emergency Measures

- 1 Oxygen in high concentration preferably 100% (by mask) to overcome anoxia. This also helps prevent a diaphragm spasm if ill.
- 2 Morphine 30-60 mg ($\frac{1}{2}$ to 1 gr) and atropine 1-2 mg ($\frac{1}{60}$ to $\frac{1}{30}$ gr) I.V. slowly and repeat every 3-4 hours. This helps venous thrombosis, general pulmonary arterial spasm that occurs.
- 3 Morphine sulfate 8-15 mg ($\frac{1}{8}$ to $\frac{1}{4}$ gr) or Meperidine Hydrochloride U.S.P. (Demerol®) 50-100 mg ($\frac{3}{4}$ to 1½ g) but not I.V. to help control pain.
- 4 If shock is present I.V. fluids should be used with great caution because of possibility of precipitating acute right heart failure.
- 5 If subsequent right heart failure develops ventilation should be performed (see page 185).

B Follow up Treatment

- 1 Observe reflexively each day for effects and adjust antithrombotic treatment if necessary.
- 2 Thoracentesis. If pleural effusion is present and makes respiration move by patient is

Prophylaxis

The prophylaxis of pulmonary infarction consists of the treatment of venous thrombosis (see page 217).

DISEASES OF THE PLEURA AND PLEURAL SPACES

ACUTE FIBRINOUS PLEURISY (WITHOUT EFFUSION)
(code No 370 100 4)

Fibrinous pleurisy is most often secondary to an underlying pulmonary disease. Whenever no apparent primary disease is present always consider tuberculosis. About 30-40% of cases of so-called primary pleurisy actually re-tuberculous. A large number of these patients develop active pulmonary tuberculosis within a 5 year period.

Diagnosis

Acute fibrinous pleurisy can usually be diagnosed mainly from history of pain in the side of the chest aggravated by respiration. The pain may be felt in the shoulder or referred to neck or abdomen in diaphragmatic pleurisy. There may be other symptoms depending on the underlying disease. The signs vary depending on the extent and nature of the lesion. Many cases have a pleural friction rub but this may be absent.

Treatment

Treatment is aimed at the underlying disease. The treatment of the pleurisy is entirely symptomatic and aimed at relieving the pain.

- A Analgesics. May be used as necessary (see page 32)
- B Ethyl chloride spray. A local anesthetic is of value (see page 126)
- C Strapping of chest with adhesive tape may afford relief by restricting movement.
- D Procain hydrochloride intercostal block may be used in more severe cases.

ACUTE FIBRINOUS PLEURISY (WITH EFFUSION)
(code No 370 100 8)

Most cases of pleurisy with effusion are secondary to pulmonary disease. However there are a small number that are primary in the pleura (hematogenous implantation of infection). Whenever sterile pleural effusion is discovered without obvious cause consider it tuberculous until proved otherwise.

Diagnosis

The diagnosis is usually relatively simple when one makes use of physical diagnosis and x-ray examination. However at times it may be impossible by these means to distinguish between fluid and thickened pleura. The case can only be differentiated by performing a diagnostic thoracentesis.

If fluid is discovered in the pleural space ascertain its nature and the presence of infecting organisms.

A Removal of Fluid for Examination

- 1 Remove 50-500 cc. Use a two-way tapcock to avoid introduction of air.

2 Inj t 50 000 100 000 units f p nicillin in 10 cc s lin i to th pl u l sp c th ough the sam n dle This is fo e ther p phyl x or actu l the apy

B Pl u l Pl id E m t on

- 1 Gr s am nat on Take sp cific grav ty to d te mune if ud t o t ns dat
- 2 Sm a and sta n f r d test on of organ sm and n t e of ell lar content C lle t a specim n in an ant o gul nt f r ll ount
- 3 Cult e on app p i te med a d inocul te guin a pig w th all fluids f m un xplain d pl ral effus o s to d m nt at p s e f tuber le b ll o fungi
- 4 Path log cal e am n t on of c ntr fug d button in s p t d ca es of mal g an y

T esting t of Po t p eumon a d Oth St i Eff s

A Spec fic M u All prophyl ct meas e ar d ect d at th primary d sea e B gun o ntanu ant b t in dosage a fo tr tm nt of pneumo a (see pag 122) until pat e t has b n feb il for 10 14 d ys r flu d is almost e tly r so bed

B G l M s re

- 1 Th r ent sis Wh ev r ach st tap is perf m d instill 50 000 100 000 un ts of peni ill n in ster le s lin a R m v dily obtai bl fl d by multi pl th r es t a at d ly nte v l if ne s ry R m val f mor than 1000 at a tim is not ad i ble b Re amin pl ur l fl d to r le out mpye n if the pl r y doe not r po d t tr atment
- 2 B d r t until p t t a feble

T e lm nt of T be c l Eff n (od N 370 123 8)

A Sp f M a s U compl at d p mary ff o a t t d t lly m m m l p l m o s y t b ul is A our e of o z d d am t lly l a d (PAS) is n d dat pto my in is m m d d (e e p g 133)

B C l M

- 1 B d t indicated as for m l m l p l m n ry t be ul i
- 2 Tho c te a R mov l of all r dly av il ble fl d (e b) ad s bl to m m el t thik d pl u a
- 3 Wh hgh f p tal g th tw we k h matogeno s dis emu tion hould b su pected

C Follow up T tment C eful follow up fo a 5 y a p i od is n s ry be m y p t nt w th p um ry t be ul s ffu ons d lo p p l m o n y t b r losis lat usu lly w th n 5 y

EMPYEMA (code No 370 100)

Empy ma u lly onda y to p l m ry i f t b t may es lt f m di ct ont m n t n f th pl r l pa thr ght n thor t is Th pat nt u lly q t ill and th o f b l and e

DISEASES OF THE PLEURA AND PLEURAL SPACES

ACUTE FIBRINOUS PLEURISY (WITHOUT EFFUSION)
(code No 370 100 4)

Fibrinous pleurisy is most often secondary to an underlying pulmonary disease. Whenever no apparent primary disease is present, always consider tuberculosis. About 30-40% of cases of so-called primary pleurisy actually are tuberculous. A large number of these patients develop active pulmonary tuberculosis within a 5 year period.

Diagnosis

Acute fibrinous pleurisy can usually be diagnosed merely from history of pain in the side of the chest aggravated by respiration. The pain may be felt in the shoulder or referred to neck or abdomen in diaphragmatic pleurisy. There may be other symptoms depending on the underlying disease. The signs vary depending on the extent and nature of the lesion. Many cases have pleuritic friction rub, but this may be absent.

Treatment

Treatment is aimed at the underlying disease. The treatment of the pleurisy is entirely symptomatic and aimed at relieving the pain.

- A Analgesic May be used as necessary (page 32)
- B Ethyl chloride spray a local anesthetic of value (see page 126)
- C Strapping of chest with adhesive tape may afford relief by restricting movement.
- D Procaine hydrochloride intercostal block may be used in more severe cases.

ACUTE FIBRINOUS PLEURISY (WITH EFFUSION)
(code No 370 100 8)

Most cases of pleurisy with effusion are secondary to pulmonary disease. However, there are a small number that are primary in the pleura (e.g. hemogenous implantation infection). Whenever sterile pleural effusion is discovered without obvious cause, consider it tuberculous until proved otherwise.

Diagnosis

The diagnosis is usually relatively simple when one makes use of physical diagnosis and x-ray examinations. However, at times it may be impossible by these means to distinguish between fluid and thickened pleura; these can only be differentiated by pneumothorax and diagnostic thoracentesis.

If fluid is discovered in the pleural space, treat it in its nature and the presence of infecting organism.

A Removal of Fluid by Examination

- 1 Remove 50-500 cc. Use a two-way stopcock attached into the tube for

SPONTANEOUS PNEUMOTHORAX (code No 306 900 5)

Spontaneous pneumothorax usually is caused by rupture of an emphysematous bleb when a undraining lung disease is present. It is also one of the complications of tuberculosis. Always investigate if tuberculosis is in a case of spontaneous pneumothorax.

Treatment

- A Bed rest until it has been largely reabsorbed. If tuberculosis is present treat accordingly.
- B Symptom treatment as indicated.
 - 1 Pleuritic pain should be treated with an analgesic at sparing doses of morphine or codeine (see page 126).
 - 2 Cough. If there is no underlying pneumonia and cough is annoying, codeine 0.015-0.06 Gm ($\frac{1}{4}$ to 1 gr) every 3-4 hours should be used.
 - 3 Aspirate if symptoms severe. If pneumothorax continues to reaccumulate in an undraining distended chest, treat with a thoracostomy (see below) may be necessary.
 - 4 Administer oxygen if dyspnea is present.
- C Surgery. In some cases of spontaneous pneumothorax when the lung does not expand or when there is a repeated episode of collapse, exploratory thoracotomy may be necessary.

TENSION PNEUMOTHORAX

(From Lung: code No 306 400 4)

(From Chest Wall: code No 377-400 4)

This is a medical emergency. The result of a valve-like action of the pleura is the escape of air from the lung or the chest into the pleural space from the outside of the lung during inspiration and cannot escape during expiration. Unless treated immediately death may result.

Treatment

A Emergency Treatment

- 1 Perform a thoracostomy immediately into the anterior part of the affected chest in the third intercostal space; aspirate the pleural space to relieve the expanding lung. The end of a rubber glove finger slit at the tip may be tied over the hub of the needle to act as a simple one-way valve. As soon as possible connect the needle to a rubber tubing and place the other end of the tube under 1-2 cm of water in a suitable container. As soon as feasible a rubber catheter introduced into the pleural space by means of a trocar should be substituted for the incising needle.
- 2 If pain is severe give morphine as follows: 8-15 mg ($\frac{1}{8}$ to $\frac{1}{4}$ g) I.V. or I.M. immediately.
- 3 Test for kinking of the tube (see page 27).

B Follow-up Treatment As for spontaneous pneumothorax (see above).

142 Pneumothorax

Treatment

- A Specific Measures Systemic administration in high dosage of appropriate chemotherapeutic or antibiotic agent as determined by examination of infecting organism. Treatment should be continued for 10 to 14 days after patient is afebrile or fluid has become sterile (see pag 514)
- B Daily thoracentesis should be performed removing as much of the purulent material as possible. Frequent physical examination of the chest with x rays must be done to avoid overlooking any loculated areas (pockets) of purulent material
- 1 Irrigate empyema cavity with 500-2000 cc saline
 - 2 Instill 100 000-200 000 units of penicillin and 0.5-1.0 Gm of streptomycin in 10 cc into the cavity at the completion of the irrigations. Continue daily until fluid in cavity has been sterile for 10-14 days or until fluid can no longer be obtained
 - 3 Various enzymatic agents have recently been prepared which digest the protein material especially the fibrin that forms in this condition. These are also introduced directly into the thoracic cavity. The principal ones available are trypsin (Trypsin[®]) and streptokinase (Streptodornase[®] (Vredas[®]). Trypsin acts by digesting all nonliving protein material. The streptokinase streptodornase attacks mainly fibrin.
- C Surgical drainage is necessary if patient does not improve in a few days or if pus becomes too thick to aspirate with a needle.

HYDROTHORAX (code No 370 522)

Hydrothorax is most generally due to congestive cardiac failure. Treatment is directed at the failure itself. In cases of respiratory embarrassment removal of the fluid is necessary.

HEMOTHORAX (code No 370 532)

Hemothorax is most commonly due to trauma. World War II experience has shown that prompt aspiration of the blood from the pleural cavity is the treatment of choice. Repeated aspirations are performed as necessary. If bleeding continues thorotomy is indicated. Great care must be taken in these aspirators to avoid possible bacterial contamination of the pleural cavity. The protolytic enzyme (see above) may be self-administered to stop the surgical removal of residual blood clot may be necessary.

PNEUMOTHORAX

Air in the pleural space can occur as a result of air entering through an opening in the chest wall (i.e. by a traumatic pneumothorax or trauma) or as a result of air entering from inside the lung.

SPONTANEOUS PNEUMOTHORAX (code No 306 900 5)

Spontaneous pneumothorax usually is caused by rupture of an emphysematous bleb when no underlying lung disease is present. It is also one of the complications of tuberculosis. Always investigate for tuberculosis in any case of spontaneous pneumothorax.

Treatment

- A Bed rest until a rash has been largely resolved. If tuberculosis is present treat accordingly.
- B Symptomatic treatment as indicated:
 - 1 Plural pain should be treated with an analgesic, aspirin or ethylhydralazine (see page 126).
 - 2 Cough. If the sound of lung pneumonia and cough is annoying, codeine 0.015-0.06 Gm ($\frac{1}{4}$ to 1 gr) every 3-4 hours should be used.
 - 3 Aspirate if symptoms are severe. If pneumothorax continues to accumulate, an underlying mediastinal shift to the left (see below) may be necessary.
 - 4 Administer oxygen if dyspnea persists.
- C Surgery. In most cases of spontaneous pneumothorax, when the lung does not expand, where there are repeated episodes of collapse, exploratory thoracotomy may be necessary.

TENSION PNEUMOTHORAX

(From Lung, code No 306-400 4)

(From Chest Wall, code No 377-400 4)

This is a medical emergency. This results from a valvelike action of the pleural surfaces of the right lung on the chest. Air is sucked into the pleural space from the trachea or from the lungs during inspiration and cannot escape during expiration. Until a stricture is made at the point of the fault.

Treatment

A Emergency Treatment

- 1 Place a 16 gauge needle into the anterior part of the affected hemithorax, 1st intercostal space, just below the axilla to the expanding lung. The end of a rubber glove finger, slit at the tip, may be inserted over the hub and if the needle is a simple one, a simple one-way valve. As soon as possible connect the needle to a rubber tubing and place the other end of the tube under 1-2 cm of water in a stable container. As soon as a free rubber catheter is introduced into the pleural space by means of a trocar, should be substituted for the needle.
- 2 If possible, give morphine sulfate 8-16 mg ($\frac{1}{8}$ to $\frac{1}{4}$ g) i.v. or i.m. immediately.
- 3 Treat shock if present (see page 27).

- B Follow-up Treatment. As for spontaneous pneumothorax (see below).

TRAUMATIC PNEUMOTHORAX (code No 377-400 3)

This is an emergency. Large open chest wounds (i.e. sucking wounds) are the most severe. The opening must be made airtight by any available means i.e. bandage handkerchief shirt or other material and closed surgically as soon as possible.

CORRECTION OF HYPOXIA AND OXYGEN THERAPY

Oxygen therapy consists of the administration of oxygen at concentrations greater than are found in the atmosphere. Increased concentrations are indicated only when hypoxia exists. The correction of hypoxia does not always require oxygen therapy. In fact in some cases of hypoxia oxygen therapy may be dangerous if not administered properly.

Oxygen therapy is always palliative. It is generally used to tide the patient over an emergency situation while the underlying cause is being corrected. Correction may not always be possible. When verbal respiration ceases, resuscitation must be instituted.

A classification of the most common causes of hypoxia and methods for their correction are found in the chart on page 146.

Dangers of Oxygen Therapy

Although much has been written about the dangers of oxygen therapy, the principal danger appears to be depression of respiration in severely hypoxic patients who have an elevation of CO_2 tension in the blood. In these individuals the respiratory center in the medulla has been anesthetized by the high CO_2 tension and respiration is under control of the chemoreceptor center's response to oxygen tension. Therefore when high concentrations of oxygen are given the chemoreceptor centers are no longer stimulated and there is a resultant decrease in pulmonary ventilation which may cause enough CO_2 retention to produce narcosis, unconsciousness, and even death. This usually occurs within a few minutes after starting high concentrations of oxygen but may occur even up to 12 hours after instituting treatment. Therefore no patient should be given oxygen unless he is under close observation during the first 30 minutes of oxygen administration and hypotension with suspected elevation of CO_2 tension should have a nurse in constant attendance and if possible some method of mechanical ventilation should be kept available.

Treatment of Hypoxia Associated With CO_2 Retention

This situation may be managed in one of two ways:

1. Oxygen may be administered by means of a automatic mechanical pressure device (e.g. body respirator Bennett apparatus) cyclically automatically or by an attendant. This is the more effective method because it promotes adequate ventilation, the CO_2 is removed more rapidly.
2. If high concentration at one's disposal cannot be immediately reduced, oxygen concentrations in the gas slowly as CO_2 is removed.

Oxygen Toxicity

Although much has been written about oxygen toxicity, there appears to be little evidence of its clinical occurrence. Many of

reported incidence of oxygen toxicity have been cases of irritation resulting from improperly humidified oxygen

TECHNIC OF ADMINISTRATION

When oxygen therapy is used proper humidification even aerosolized water or saline must be maintained

OXYGEN AT ATMOSPHERIC PRESSURE

Oxygen is most commonly administered at atmospheric pressure. It is indicated when hypoxia can be controlled adequately by the means alone. For indications see page 146.

Various methods of administering oxygen at atmospheric pressure are in use. Below are tabulated those methods which are most commonly used with adults and the oxygen concentrations which can be achieved.

Method	Usual Oxygen Concentration	Usual Rate of Oxygen Flow (L./min)
Tent	40-50%	Flow 15-20 Minimum at 12-15
Catheter		
a Nasopharyngeal (metallo rubber)	20-40%	6-8
b Oropharyngeal	30-40%	6-8
Mask		
a BLB or equivalent	80-100%	8-10
b Expendable plastic mask	40-60%	10-15
c OEM or Bennett face mask	80-100%	6-8

Oxygen Tent

A Advantages

1. Gives moderate concentrations of oxygen at maximum comfort to the patient.
2. Can be used with ill and uncooperative patients.

B Disadvantages

1. Must be kept very dry and to operate at
2. Cannot achieve high concentrations of oxygen.
3. If not operated properly oxygen concentration can anaerobically fall and carbon dioxide is likely to accumulate.

Nasal Catheter

This apparatus consists of a uethal tent (French No. 10-12) with 4-6 small holes in the terminal inch a reduction valve mechanism and a humidifier bottle. A bilateral plastic or metal anastomosis extends about 1 inch into the trachea and is a suitable

A Technique

1. The tent should be lubricated with petroleum and placed every 6-12 hours.

HYPOXIA COMMON CAUSES AND METHODS OF CORRECTION

Physiologi al Classification	Clini al Conditi	Treatm t
Hypoxia With Normal Lungs Deficient atmospheric oxygen	High altitude flying	Oxygen at atmospheric pressure
Airway obstruction	Gas ulcers (tissue anoxia) Foreign bodies Atelectasis Mucous plugs Paralysis of vocal cords Epiglottitis Tongue in mouth	Remove cause Clear the airway Oxygen usually needed Resuscitate
Paralysis of respiratory muscles	Anthrax Poliomyelitis	Respiration
Paralysis of respiratory center	Encephalitis Myelitis	Respiration

Hypoxia With Abnormal Lungs Diminished efficiency of functioning alveoli	Pneumonia	Oxygen at atmospheric pressure
	Emphysema (acute)	Oxygen by intermittent positive pressure atmospheric pressure
Improper mixing penetration of inspired gases	Emphysema bronchial asthma	Oxygen by intermittent positive pressure
Impaired diffusion as in emphysema membranes	Pulmonary edema	Oxygen by intermittent positive pressure (in an emergency position)
Valvular Arterial Shunt	Congenital heart disease	Oxygen in hypofluoride fluid preparation

Hypoxia Due to Inadequate Oxygen Transport by Blood		
Diminished effective hemoglobin	Anemia CO poisoning Methemoglobinemia	Correction Oxygen at atmospheric pressure Artificial CO-Absorbing fluid
Impaired utilization	Cyanosis poisoning by drugs Shock	Oxygen at atmospheric pressure If pulmonary edema positive pressure Oxygen at atmospheric pressure positive pressure and glucose to electrolyte balance in shock

Hypoxia Due to Impaired Tissue Oxygenation	Circulatory poisoning by drugs (cyanide)	Resuscitation Oxygen at atmospheric pressure Appropriate fluid (page 538 & 540)
--------------------------------------------	------------------------------------------------	------------------------------------------------------------------------------------------

- 2 It may be placed in the nasopharynx or inserted into the nostril but concentration is usually only about 25-30%
 - 3 Piece in oropharynx for concentrations up to 40%. To calculate the approximate distance the tube must be inserted, measure the distance from the external nares to the tip of one ear lobe using the tube to measure with. Then pass the tube through the nose into the oropharynx. When the patient begins to swallow withdraw the tube about $\frac{1}{2}$ inch and secure it in position.
 - 4 The binasal cannula gives concentrations about the same achieved with the nasal cannula.
- B Advantages** The nasal cannula is the cheapest method of administering oxygen and is more comfortable than a mask.
- C Disadvantage** Very high concentrations of oxygen are not obtainable and mucosa may dry with ordinary humidification.

Masks

- A Apparatus**
- 1 BLB masks. Nasal or oronasal rebreathing mask with rebreathing bag. The disadvantage of this mask is that with low flow of oxygen (under 6 litres per minute) CO_2 tends to accumulate. The face may also be reluctant to inspiration from flat rebreathing bag.
 - Expended plastic masks. Require high oxygen flow. Low oxygen concentration achieved.
 - 3 OEM and Bennett face masks. Similar to BLB mask but do not permit rebreathing. In bag utilizes flutter type valve so rebreathing of CO_2 is not possible.
- B Advantages of Masks**
- 1 High to 100% concentrations of oxygen obtainable without the use of pumps (except for plastic masks).
 - 2 Both OEM and Bennett masks have adjustable settings so that oxygen concentration can be varied from 50 to 100%.
- C Disadvantage** Tight fitting masks cannot be tolerated by a patient.

OXYGEN UNDER PRESSURE

Various positive pressure breathing devices have been developed which allow oxygen to be administered under slight positive pressure during the inspiratory phase. Although originally the devices were employed for resuscitation (usually with a positive pressure phase in expiration) the value of intermittent positive pressure in the treatment of various acute and chronic pulmonary and cardiac conditions was soon recognized.

Physiological Effect

The principal physiological effect of oxygen administered by positive pressure is as follows:

- 1 Helps overcome resistance to gas flow and widens the bronchioles permitting more efficient cough and bronchial drainage.
- 2 Increases intrapulmonary mixing creating more uniform alveolar ventilation.

- 3 Decreases residual volume
- 4 Inhibits fluid extravasation into the alveoli (when use of aid in pulmonary edema)
- 5 Interferes with venous return to the right heart with consequent decrease in cardiac output and blood supply to the lungs. This latter phenomenon is of value in management of congestive failure especially with associated pulmonary edema. In shock on the other hand it is a disadvantage and often contra-indicates the use of positive pressure devices in this condition.

PRINCIPAL METHODS OF POSITIVE PRESSURE BREATHING

Method	Pressures During Respiration	Indications and Uses	Remarks
Mouth to mouth or mouth to endotracheal tube	Positive pressure in inspiration	Resuscitation especially useful with children and newborn infants	Most primitive method of positive pressure but may be very effective. Oxygen administration at lower than atmospheric concentrations.
Bennett positive pressure therapy unit (motor or oxygen powered)	Positive pressure in inspiration. May use oxygen, air or oxygen helium mixture.	Mainly for therapy of chronic pulmonary diseases. Also useful in pulmonary edema.	Interferes with venous return to right heart so contraindicated in forward failure. Especially useful in hypoxia due to improper mixing of gases and for pushing oxygen across impaired membranes (see page 149).
Oxygen injector mask (Barach) metered for positive pressure	Positive pressure in expiration. Used with oxygen.	Advocated for pulmonary edema.	Least efficient. Positive pressure applied at wrong place in respiratory cycle to be of benefit. Also very tiring to breathe against resistance.
Commercial resuscitators of the suck and blow type: Stephenson, Emerson, E and J, etc.	Positive pressure in inspiration and negative pressure in expiration. Generally employ oxygen.	Resuscitation.	Most effective means of resuscitation. Let interfere with cardiovascular dynamics although the usual pressure relationship in inspiration and expiration is reversed. Negative phase may cause pulmonary edema in predisposing condition.
Kreiselman hand bellows or Emerson bellows	Positive in inspiration.	Resuscitation.	Inexpensive resuscitator useful mainly when more expensive apparatus are not available.

Bennett Positive Psychology Unit

The Bennett is one of the most effective of the available
pre-breathing devices. It may be used with an intermittent
nebulizer or with Mist-O₂ Co.® titanium nebulizer for good humid
if caution is observed in administration of various anesthetic volatile
and sedatives (lowering agents). It is particularly useful way
to administer aerosols for treatment of the terminal
bronchus and alveoli. Excellent instructions are supplied with
the unit. A popalappatus also includes that cycles automatically
Clinical indication and use as follows:

- 1 B ch i l stum b p e ally w th b r nchod l i r u e f l
mainly in th acut att ck
- 2 Chr nic mphy em id opath c or ac ompanying f b s s
p um o o is et B st res it app rently whe
bro chodilat s e used Mu t b sed aut usly or w th
ut mati cv lung in patie t w th s v re hypoxia nd ele
v ted CO₂ tensi n (S dan ers of oxyg n th apy p ge
144) In th conditio s th rapy m t b employ d 2 4
t m d ly fo bout 20 min te per tr tm t T e tme t
gl in o es f 5 30 d ys whi h m y b ep at d as
dicat d
- 3 Bron h t s A fo mphy m ab Ant b ot cs by
os ia oft n ful n th s c nd t on
- 4 Pulmo ary ed ma Esp ci lly u f l wh n s oc ted w th
s e a ox M st b s d w th gre t ut on if sh k
(f rwa d fal) is pres nt
- 5 I r t ting g s and fum s V y lu ble spe ially with
any ted pulmon y dema U e u t i lungs h ve
les ed
- 6 Atel tas Se p ge 137
- 7 R p tory d p o M st be u d w th ut on f
culato y f l re als p se t
- 8 Right hea t f ilur Helps t hypo r l ve b r d n
n r ght hea t Ex ll t i n n g m nt f t ght he t
fa lu in orjunct w th th m s r (se pag 181)

PRINCIPAL METHODS OF PRESSURE ALTERATIONS TO THE CHEST WALL

Method	Pressures During Respiration	Indications and Uses	Remarks
Artificial respiration	The best method to use is to pull on arms to expand chest for inspiration pressure for expiration (see below)	All respiratory failures when no other method is available	Entirely physiological when proper method is used (see page 151)
Body respirator	Negative pressure (suction) for inspiration usually used. May use positive pressure (expiration) in attempt to overcome venous pooling	Respiratory failure when prolonged aid is needed	Since negative pressure applied to entire body may diminish cardiac filling due to pooling of blood in extremities and trunk may be dangerous in forward failure
Cuirass respirator		As for body respirator especially in convalescent or stabilized poliomyelitis	More physiological than body respirator. Less interference with circulation. Often uncomfortable or hard to fit for prolonged time. Unsatisfactory in severely poliomyelitis
Rocking bed	Respiration controlled mainly by abdominal contents dropping away from or pushing diaphragm	Less degrees of respiratory failure as above	Momentarily excellent for improving circulation and respiratory dynamics and respiration in patient with combined respiratory and body paralysis

ARTIFICIAL RESPIRATION

Artificial respiration must be administered promptly to persons whose respirations have ceased whether due to drowning, suffocation, electric shock, or other accidental cause. *Manual artificial respiration should never be postponed while waiting for the arrival or institution of treatment with a mechanical resuscitator.*

This procedure replaces spontaneous respiration and provides oxygen to the tissues until the paralyzed respiratory center recovers and resumes its normal function. As long as the heart continues to beat, the patient has a chance of recovery, and this may occur even after many hours of artificial respiration.

The push-pull methods of artificial respiration are more than twice as effective as the simple push methods (e.g., Schafer)

The generally approved method at present is the arm lift-back pressure (Nielsen) the hip lift-back pressure and the arm lift-chest pressure (Sylvester) methods are less efficient than the Nielsen method but are preferable to the simple push methods

General Procedure

- A *Clear the airway and begin artificial respiration at once and lay of only a minute or two reduces the victim's opportunity for recovery*
- B *Do not stop artificial respiration until normal respiration is established or until rigor mortis begins*

Technic of the Arm Lift-Back Pressure Method (Nielsen)

- A Position of Patient The patient is placed prone with his head turned to one side and resting on the backs of his hands
- B Position of Operator The operator kneels (on either or both knees) at the patient's head and then grasps the patient's arms at a point between the elbows and the shoulders
- C The Rate of Resuscitation The rate of resuscitation is maintained by 10 to 12 complete cycles a minute. This rate can be timed by a watch or by counting the following numbers 1001 1002 1003 1004 1005 1006 and the repeating. This manner of counting requires about 1 second for each number. These seconds should be allowed for each arm lift and for the



1 Place hand for arm lift



2 Rock backward and lift arms



3 Place hand for back pressure



4 Rock forward and press back

The Arm Lift Back Pressure Method of Artificial Respiration

back pressure. When possible, operations should be alternated at 20 to 60 minute intervals.

D Procedure of Resuscitation

- 1 Arm lift. The operator lifts the patient's arms upward and toward himself as he rocks backward on his knees. The arms of the operator are kept straight during the entire procedure. This arm lift enlarges the thoracic cage and causes inspiration. The arm lift is continued until resistance is met; the patient's arms are then returned to the ground and the operator places the palms of his hands on the patient's back.
- 2 Back pressure. With the palms of the hands on the lower part of the shoulder blades and the fingers extended over the thoracic cage, the operator rocks forward on his knees and with his arms straight exerts hard pressure directly downward on the thorax until resistance is met. The cycle is then repeated.

Mechanical Resuscitators

In competent hands, mechanical resuscitators are more effective than and should replace manual artificial respiration as soon as available at the site of emergency. However, it should be emphasized that a mechanical resuscitator should only be used by trained personnel and when it is in proper mechanical condition.

AEROSOL THERAPY

There are two types of aerosol therapy: intermittent and continuous. Intermittent therapy is the more commonly used. Recent work suggests that continuous administration of water or saline by mist of fine particle size allows for better humidification with less irritation from oxygen and appears to be more physiologically than steam inhalations. Continuous aerosol should probably be employed in all conditions where there is a potential for bronchial irritation. Surface tension lowering agents, antibiotics, and bronchodilators may be used by this method.

The administration of antibiotic agents by aerosol inhalation has been of value in some lung infections. Certain pieces of equipment are necessary in order to administer aerosol therapy.

- A Nebulizers producing particles smaller than 5 to 10 μ in diameter. The most satisfactory nebulizers are the Vaponephrine® model and the DeVilbiss No. 40®. For continuous administration of aerosols, apparatuses with large capacities are available (e.g., Mist-O-Gen®, Humidox®).

B Sources of Pressure for Nebulizing Drug

- 1 Oxygen from a cylinder at 6-10 liters per minute is usually used.
- 2 Compressed air from a diaphragm type compressor [Caution: Do not use an oil-sealed pump.]
- 3 Manual pumping devices (e.g., foot bellows or pumps) have been employed but are generally not very useful as they tend to tire the patient and are inefficient.
- 4 Nebulizer with hand bulb may be employed but is quite inefficient because it does not produce an adequate flow for a sufficient length of time.

C Drug and C ent tions Employ d (Should b prepa d f sh daily) Th f que y and d at n f r atme t dep nds upon the d eas and t s v rity

1 Antib otics

- a Pen illin Usual dose is 50 000 100 000 units pe treatme t Dilute in 1 0 2 0 cc of w t r
- b St eptomycin 0 25 0 5 Gm in 1 0 2 0 cc of wat r Oxytetra yclin (T r amy n[®]) a rosol 50 100 mg in 1 0 2 0 c 75% propylene glycol

2 En ym

Although tryp in (Trypt r[®]) has b advocat d to d lve th k ten ous m us or dead t su in chronic b onchitis or b onch ctia i untowa dr ctions and b arre changes in cell h v been ob erv d with ts u It must be us d with gr at uti n following instructio s car f lly Do ge 1 3 c f trypsin ol t n (40 000 un ts/) prep ed by dissolving th d ypowd in a sp al buff r (pH 7 1) Administ 4 times d ily f up to 4 5 days f r ach o e D ot employ w thin one week after f ank b m rh g My be ombin d with p icillin t ptomy in and bron h l dil tors

- b Oth ym h v b e u d [g des xy bo ucl s (Dorna e[®])] b t mu t st ll b on der d expe im ntal

3 Br nchodil t rs

- a I p ot r ol (Isup l[®] Al drin[®]) 0 1 0 5 f l 100 or 1 200 ol t n
- b Epin phrine (adr nal) 0 5 c f l 100 lut on

- 4 S f c t ns on lowe ing ag nts V ous rf e ten ion l w r gag nts hav be adva d to s d in sp e ding and f mation of osols Th r valu s till uncertain Am ng the dr gs e Al vair[®] ethyl al h l tc

D M th ds f Adm ni t ation

- 1 Oral nhai t u d ng inspi t ry ph F th gre t st ff t d ffic n y th a r ol ho ld be inhal d th ough the mouth

Co ti vou p s f m xyg n tank A Y tub s rt d b tw en b h r and ur of p ss re Nebul z t n w ll c u o ly wh n th attach d e d of th Y tub s l ed by the thumb a f g the e lly a f w cond d lay befor th e ol ar v at the mouth p

- b Int mitt t p ss e (g foot b llow p mp) is pplid du ing inspi at n

- 2 If th p t nt is u abl to op te th b h r may be s d w th an xyg n m sk wh h b s a b thng bag at t h d Th n b l pla d b tw n ma k and oxyg n

Chapter 7

DISEASES OF THE HEART*

CONGENITAL HEART DISEASE

Congenital disorders of the heart which are amenable to surgical correction are presented below

Although in many instances the diagnosis of congenital heart disease can be made on clinical grounds alone, most cases require special studies which are best performed in medical centers. Such procedures as cardiac catheterization, venous and retrograde aortography, and tomography are not suitable for the occasional investigation; they are difficult to perform and interpret and require skilled teamwork.

Curative intracardiac surgery with a pump-oxygenator is now feasible with a reasonable mortality risk in a few medical centers. It is the procedure of choice for tetralogy of Fallot, ventricular septal defect, pulmonary stenosis with normal aortic root, atrial septal defect with an ostium primum defect, and congenital aortic stenosis. Not all patients with these conditions require reparative surgery, however.

Tetralogy of Fallot (code No. 413.0x0)

All children with the tetralogy of Fallot should be operated upon because so few reach the age of 21 if untreated. Syncope is an urgent indication for operation. Because of the higher operative mortality rate in the adult age group, patients over the age of 21 are operated upon only if they are seriously disabled.

Pulmonary Stenosis With Normal Aortic Root (code No. 413.0x0)

This condition is frequently considered a slowly recognizable being one of the most common congenital lesions. Depending upon the severity of the lesion, the patient may be asymptomatic or may show all degrees of cardiac disability up to severe cardiac failure with low cardiac output.

If severe stenosis is manifested (right ventricular systolic pressure of at least 100 mm Hg and progressive right ventricular enlargement), surgical treatment is indicated.

If the stenosis is mild, the patient may be asymptomatic for years.

For drugs used in treatment of cardiovascular diseases
pages 195 to 206

Transposition of the Aorta (code No. 452.017)

The condition is recognized by the combination of a centrally located congenital lesion, a dominant α wave in the venous pulse and evidence of left ventricular hypertrophy clinically and electrocardiographically. A Blalock anastomotic operation is the treatment of choice.

Patent Ductus Arterialis (code No. 40.000)

This relatively common lesion arises secondary from complete absence of symptomatic functional failure.

The indication for ligation of a patent ductus arteriosus in the presence of pulmonary hypertension should not be established. Current opinion favors ligation whenever flow through the duct is predominantly or intermittently from left to right.

Because of the low operative mortality rate (less than 1%) in killed animal studies on a dog series is recommended in all individuals under the age of 20 perhaps 30. The mortality rate also becomes higher as the patient becomes old. This recommendation is in recommending surgery in adults who are asymptomatic and have no left ventricular hypertrophy. Subacute bacterial endocarditis is the major hazard in this group.

Coarctation of the Aorta (code No. 461.018)

The condition is characterized by limitation of the aortic blood pressure in the arms but not in the legs, a weak distal femoral pulse, a prominent thoracic aortic arch, diminished peripheral pulses, a systolic murmur heard at the base of the heart anteriorly, diastolic and signs of late arteriosclerosis, return of the constricted aorta. If the distal ends of the thoracic aorta are visible, the diagnosis is established by angiography or aortography.

Reconstruction of the constricted aorta of mild to moderate degree is the indication for the patient due to aortic coarctation and the surgical mortality is in the neighborhood of 3% even in the best hands. For this reason not all physicians recommend routine resection in symptomatic individuals. The risks of the disease are such however that if a killed dog logical surgery is a suitable alternative up to the age of 20 years should be considered. Between the age of 20 and 35 surgery is advisable if we are to make it clear that the patient is doing badly.

Atrial Septal Defect (code No. 41.0xx)

Mortality rates of atrial septal defects do not require surgery. Surgery should be withheld in severe lesions with pulmonary hypertension and associated shunt because of the risk of cerebrovascular thrombosis with large left to right shunt (more than 2 to 3 times a systemic flow) without evidence of pulmonary arterial resistance should be operated upon. The exact procedure of choice has been established by the following factors:

1. Established by the following factors: a. If the defect has been complicated with atrial fibrillation, it is usually necessary to perform a complete atrial septectomy.

Accessory Pulmonary Vessel (code No. 488.02)

Surgical removal of total anomalous pulmonary venous drainage is now possible with safety by passing a catheter.

Ventricular Septal Defect (cod No 413 0xx)

Ventricular septal defect vary in severity from trivial asymptomatic lesions with normal cardiac hemodynamics to extensive lesions causing death from cardiac failure in infancy. The former do not require surgery. The ideal case for curative repair with cardiac bypass techniques is one with a large left to right shunt, left ventricular hypertrophy, and only moderate pulmonary hypertension. When severe pulmonary hypertension is present (pulmonary arterial pressures of more than 85 mm Hg) and the left to right shunt is small, the surgical mortality risk is about 50%. If the shunt is reversed surgery is contraindicated.

HYPERTENSIVE CARDIOVASCULAR DISEASE

(code No 400 533)

Hypertension per se is a manifestation, a hemodynamic sign, and the course of the disease is adversely affected by:

1. Cardiac failure secondary to increased work of the heart and relative or absolute coronary insufficiency
2. The development of atheromata, especially in the cerebral and coronary arteries with syndromes resulting from vascular occlusion
3. Acute vascular necrosis resulting from rapid sustained rises of diastolic blood pressure usually exceeding 130 mm Hg which produce the complication known as the malignant phase. Renal failure occurs almost exclusively in this last group.

Evaluation of the Hypertensive Patient

Hypertension is a nonspecific sign and may be present in a variety of diseases, some of which are curable or can be modified by treatment. The first step in the treatment of patients with elevated diastolic blood pressure is to recognize correctable reversible conditions in which hypertension occurs. These diseases include unilateral or bilateral kidney follow-glandular disease, endocrine disorders such as pheochromocytoma, Cushing's disease, vascular disease such as coarctation of the aorta, and acute glomerular nephritis.

The severity of the vascular hypertension and the integrity of the vital organs commonly affected by hypertension (heart, brain, fundi, kidney) must be assessed before the appropriate treatment be planned.

Classification of Severity of Hypertension and Its Complications

Hypertension may be classified as follows:

- Severe Papilledema or exudates in the fundi, cardiac failure or disabling dyspnea, discharging coronary insufficiency, repeated cerebral thrombosis with neurological sequelae, rapidly developing diastolic hypertension with progressive left ventricular hypertrophy.
- Moderate Signs of left ventricular hypertrophy, arteriosclerotic changes in the fundi, old cerebral thrombosis with sequelae, easily controlled coronary insufficiency.
- Mild Diastolic blood pressure below 125 mm Hg with minimal or no objective signs of vascular damage in fundi, heart, brain, or kidney.

Method: Atrial Low Ring Blood Pressure

- A Dose (See below) R w ifia ompo nds v atrum com pound hyd alaz e hydro chloride (Ap esol[®]) m lho um ompo nds and m camylamin (I si e[®]) and thiocyanat s z dnt ite Chloroth zide (Duril[®]) i an wor l d retic which n bout 50% of ses redu s the dose requi d of ga glo c bio ki gag nts th vailabl d ta e sca ty D r l[®] dos g 0 5 0 75 Gm (7 1/2 12 gr) a d y in d ded do es ha b en us d with due ca tion for elect olyte d plet on
- B Sympathet my
- C L w it r ce d i t
- D Othe (g psy hotherapy s d ton)

Indications for Potent Hypotensive Drugs and/or Sympathet my

- A D f i t i d t i
- 1 Malign t hype te io
 - 2 Hyp rt s e rdia fail re wh n a t myo di l inf rc t on ha b n xclud d (f p s ble)
 - 3 Rapidly advan g dia tolic blood p ess e with l ft e t c la hyp rt phy and d l t ti n Evid c of d teriorat n in th h t nd fundi (udate d hemor hage) sp c ally in young (p ticula ly m l) indi duals
- B P ob bl i d t s (Explo to y St g s)
- 1 R cur e t mild ce bral th ombos with n u l gi l e q l e d high di stoli pr s e i t ct bl na y ns ffs ie y nd h gh dia tolic pr su
 - 3 A ymptom t m n with di stol blood p es s betw 125 and 130 mm Hg b t without other evidence of complica tion of hyp t sion
 - 4 Se e e i tractabl hyp t h d ch s n the ab n e f ob us emotional t s
- C N t i d t d W th Pr s nt Kn wledg
- 1 Mld be gnes e t l hyp rt i m ddle ged wom n w th ut object e d e of va ul det r at on or c m pl at o
 - 2 E rly t s nt hyperten io in young i d id l s w thout obje ti e ide c of va cul d t io t on o c m pl to

Hypot i D gs

M nyp t is w th hyp te sion e p ally middle aged women live m y years n mfort The ef great ca e hould be e et is d a d l d to of e significant dec e s in th spa of lif obta d b fo e ubje t g these p ti nt to the diag e bl side effe ts d p t ntial dang of contin sp g am of d ug the py Hype te on ari st aki gly i sev rity i diff re t p ti ts t e tm t at p nt should b va i d dep di g n th s ity of the hyp t s on nd the p n of m pl tion

Se id g n w vailabl b t th i d at s f r th i e ull t l a ly d fi d P t t with moderat or ev f m f th die s h uld b gi e th benefit of a trial of th r py Us th l a t t x d g fo mild hyp rt si Ov s p r i d of m ths y alight to mod t low ring of the blood p s u may pr v to d c s o possibly e e s e the va ul r m pl i tions f hyperten ion C mbinati f drug m y p o t be s f l b t h y a diff ult to e aluat R w olf a i s h b t t le t d th ompli tio s f m its us a e l e s t

In most severe cases ganglionic blocking agents should be considered but in some instances of less severe disease it will be worthwhile to begin with rauwolfia and then either add another drug or change to a more potent drug if rauwolfia is not effective

- A Rauwolfia Drugs** Rauwolfia has a relatively slight hypotensive action but may be useful because of its mild sedative effect and its value as an adjunct when combined with methonium compounds veratrum or hydralazine (Apresoline®). Although it is the least toxic of the hypotensive drugs nasal stuffiness may be annoying. Gastric hyperacidity may occur with larger doses. Sodium retention or severe depression may occur in which case the drug should be withdrawn.
- Dosage**
- 1 Reserpine N N D (Reserpid® Serpasid®) 0.1 to 0.25 mg (1/800 to 1/250 gr) tid orally at onset. Maintenance therapy 0.25 to 0.50 mg/day
 - 2 Rauwolfia N N D (Raudixin®) (Rauwolfia serpentina whole root) 100 to 200 mg (1 1/2 to 3 gr) daily
- B Veratrum Compounds** These compounds have not received universal favor because of the narrow margin between their therapeutic and toxic effects nausea vomiting and weakness. More recently purified preparations particularly Protoveratrine A and B N N D have been found useful especially in hypertensive emergencies. In these situations (heart failure complicating acute nephritis the convulsions of eclampsia hypertensive pulmonary edema) give protoveratrine as follows
- 1 Acute hypertension
 - a I V 1.5 to 1.8 mg/Kg. The hypotensive effect lasts 1 to 3 hours
 - b I M 1.2 mcg/Kg every 8 hours
 - 2 Chronic hypertension 0.4 to 1.5 mg (1/150 to 1/40 gr) orally tid or qid after meals (average dose). The dose must be carefully regulated at times a difference as slight as 0.5 mg may make the difference between effective vomiting or the absence of toxic symptoms
- C Hydralazine Hydrochloride N N D (Aprisolin®)** The initial dosage of this drug is 25 mg (3/8 gr) orally tid progressively increasing to a total dosage of 300 mg (5 gr) a day. The results of the oral use of this drug as a sole method of therapy are often not impressive but some patients obtain a hypotensive effect. Because Aprisolin® is the only hypotensive agent which increases the renal blood flow it may be useful as an adjunct to oral methonium compounds (ganglionic blocking agents).
- Toxic side effects are common. The most important are headache and palpitations with tachycardia. A syndrome resembling diffuse collagen disease has occurred usually after large doses have been used for many months.
- D Ganglionic Blocking Agent** The most frequently used of the agent is Prazosin mesylate N N D (Apizyn®). Chlorisondamine Hydrochloride N N D (Eccolad®) and Mecamylamine Hydrochloride N N D (Iversal®). They can be used orally subcutaneously or intravenously. At the present time with the exception of Iversal® the oral route has the disadvantage of small and irregular absorption from the gastrointestinal tract with resultant unpredictable falls in blood pressure.

Basic principles

- a Hospitalize patient under close supervision
- b Start with small initial dose and increase gradually depending upon the tolerance and response of the patient. The degree of reduction of pressure should be only moderate in the first week or so and no attempt should be made to reduce the pressure to normal until it has been demonstrated that the patient can tolerate systolic pressures of below 160 mm. Hg without hypotensive symptoms
- d Postural hypotension which is great at the height of the effect of the drug should be considered not only as a potential danger to the patient but also as a therapeutic weapon to prolong the hypotensive action of the drug after the peak effect has worn off

- 2 Oral ganglionic blocking agent. A trial of 2 or 3 weeks usually required before the dose required to lower the blood pressure to a level approximating 160/100 mm Hg can be determined. The patient may then be seen as an out patient and the dose gradually increased to that level which produces the desired fall of pressure. Whether the desired level of pressure at the time of peak action of the drug is in the range of 150 to 160 mm Hg systolic or whether it is slightly higher results in mild hypotensive symptoms on standing has not been determined. Constipation is to be avoided in patients receiving ganglionic compounds because it increases the absorption of the drug. Laxatives should be given to ensure a daily bowel movement.

Although the determination of the proper drug dosage is difficult, it is usually decided satisfactorily if taking diastolic pressure of 100 mm Hg or less as a guide. Since the effectiveness of the drug cannot be determined by usual blood pressure readings in the physician's office, other methods have been used to determine effective dosage.

- (1) Home blood pressure readings by either the patient or a responsible member of the family are recorded and when to the physician at his regular visits. On the basis of these readings the physician may increase or decrease the dose. The patient is instructed to decrease the dose whenever the blood pressure falls below 150/90 and not to take a dose if the blood pressure is below 130/80 in the recumbent position.
- (2) Motional sensitivity test for one minute prior to a dose is added to prevent excessive hypotension. If an individual can stand motionless for one minute prior to a dose, his blood pressure will be sufficiently high so that an additional dose of the drug can be taken. This only guard against excessive dosage and does not indicate when the dose has been inadequate.
- (3) Prolonged hospitalization for a day or two to determine basal blood pressure readings. These readings are often 50 to 100 mm Hg less than casual readings obtained in the doctor's office and can be used to control the dosage of the ganglionic compounds.

The initial dosage of the ganglionic blocking compounds orally is as follows:

- a Hexamethonium 125 mg (2 gr)
- b Prolinium Tartrate N N D (Ansolyse[®]) 10-20 mg (46 1/3 gr)

In most severe cases ganglionic blocking agents should be considered but in some instances of less severe disease it will be worthwhile to begin with rauwolfia and then either add another drug or change to a more potent drug if rauwolfia is not effective.

A Rauwolfia Drugs Rauwolfia has a relatively slight hypotensive action but may be useful because of its mild sedative effect and its value as an adjunct when combined with methonium compounds veratrum or hydralazine (Apresoline®). Although it is the least toxic of the hypotensive drugs nasal stuffiness may be annoying. Gastric hyperacidity may occur with larger doses. Sodium retention or severe depression may occur in which case the drug should be withdrawn. Dosage:

1 Reserpine N N D (Reserpoid® Serpasil®) 0.1 to 0.25 mg (1/500 to 1/250 gr) t.i.d. or b.i.d. at onset. Maintenance therapy 0.25 to 0.50 mg/day.

2 Rauwolfia N N D (Raudixin®) (Rauwolfia serpentina whole root) 100 to 200 mg (1 1/2 to 3 gr) daily.

B Veratrum Compounds These compounds have not received universal favor because of the narrow margin between their therapeutic and toxic effects nausea vomiting and weakness. More recently purified preparations particularly Protoveratrine A and B N N D have been found useful especially in hypertensive emergencies. In these situations (heart failure complicating acute nephritis the convulsions of eclampsia hypertensive pulmonary edema) give protoveratrine as follows:

1 Acute hypertension

a I.V. 1.5 to 1.8 mcg/Kg. The hypotensive effect lasts 1 to 3 hours.

b I.M. 1.2 mcg/Kg every 8 hours.

2 Chronic hypertension 0.4 to 1.5 mg (1/150 to 1/40 gr) orally t.i.d. or q.i.d. after meals (average dose). The dose must be carefully regulated at times a difference as slight as 0.5 mg may make the difference between sudden vomiting or the absence of toxic symptoms.

C Hydralazine Hydrochloride N N D (Apresolin®) The initial dose of this drug is 25 mg (3/8 gr) orally t.i.d. progressively increasing to a total dosage of 300 mg (5 gr) a day. The results of the oral use of this drug as a sole method of therapy are often not impressive but some patients obtain a hypotensive effect. Because Apresoline® is the only hypotensive agent which increases the renal blood flow it may be useful as an adjunct to oral methonium compounds (ganglionic blocking agents).

Toxic side effects are common. The most important are headache and palpitations with tachycardia. A syndrome resembling diffuse collagen disease has occurred usually after large doses have been used for many months.

D Ganglionic Blocking Agents The most frequently used of these agents are Pentolinum Tartrate N N D (Apodyn®) Chlorisondamine Chloride N N D (Ergolid®) and Mecamylamine Hydrochloride N N D (Inversine®). They can be used orally subcutaneously or intravenously. At the present time with the exception of Inversine® the oral route has the disadvantage of small and irregular absorption from the gastrointestinal tract with resultant unpredictable falls in blood pressure.

- a Acute hypotensive reactions are manifested by faintness, weakness, and nausea and vomiting. The patient should be instructed to lie down immediately when these occur and place his feet higher than his head. Unless the hypotensive effect is too severe, the symptoms pass off rapidly with this positional assistance. If the symptoms persist, give a vasopressor drug such as Phenylephrine Hydrochloride U.S.P. (Neo-Synephrine®) or Methoxamine Hydrochloride U.S.P. (Vasoxyl®) subcutaneously as a low concentration intravenous infusion of Levaterol Bitartrate U.S.P. (Levophed®) 4 mg/lite (see page 419).
- b Aute or progressive renal failure does not decrease the blood flow or filtration pressure markedly and requires discontinuance of the drug.
- c Vascular thromboses are a hazard in older patients who suffer severe hypotensive falls.
- d A low sodium diet potentiates the action of methonium compound, and if an individual is receiving fixed doses of the drug, a gradual decrease in sodium diet hypotensive symptoms may occur. It is usually desirable to place the patient on a 1.5 Gm (22 g) sodium diet at the onset of therapy.
- e Alcohol, hot climate, vasodilator drug, vigorous exercise, and salt depletion potentiate the action of methonium compound.
- f Parasympathetic effects (due to parasympathetic blocking): Blurring of vision, constipation, dryness of the mouth can be counteracted in part by the use of neostigmine orally in doses of 7.5 to 15 mg ($1/8$ to $1/4$ g).

Surgical Procedures

- A Sympathectomy. The therapeutic value of sympathectomy has been highly controversial although many authorities agree that it prolongs life when directed on patients with early malignant hypertension when renal function is good.
- B Adrenalectomy. These latter findings indicate that have not been impressive although meprits with severe hypertension significantly reduced the deleterious effects.

Low sodium diet

A rigid low sodium diet containing 350 mg ($5\frac{3}{4}$ gr) of sodium per day effectively suppresses but the diet is also a diuretic and the patient is not able to eat the food that he and his family like. It would not be indicated only in the treatment of hypertension so which drug therapy is still debated. The diet should also be used as an adjunct to the treatment of hypertension with drug. Food table for the low sodium diet page 55.

Psychotherapy

Clinical evidence is available to indicate that the hypertension is partly of the emotional conflict part of the so-called psychoneurosis and is partly of the endocrine and endocrine. Emotional disturbance is a factor in the hypertension and is a factor in the hypertension and is a factor in the hypertension.

- c Chlorisondamine Chloride N N D (Ecolid[®]) 10 20 mg
($\frac{1}{6}$ $\frac{1}{3}$ gr)
- d Mecamylamine Hydrochlorid N N D (Inversine[®])
1 2 5 mg ($\frac{1}{60}$ $\frac{1}{24}$ gr)

3 Parenteral ganglionic blocking agents

- a Hexamethonium ion The initial dose is usually 2 5 5 mg
($\frac{1}{24}$ $\frac{1}{12}$ gr) of the ion given subcutaneously If no untoward effect occurs the dose can be repeated in 12 hours On the second day 5 mg ($\frac{1}{12}$ gr) may be given twice at 12 hour intervals and the dose gradually increased On discharge from the hospital in 2 to 3 weeks the average patient receives about 75 mg ($1\frac{1}{4}$ gr) twice daily In some patients it may be necessary to give the drug 3 times a day but the increase in dosage should always be made gradually (2 5 5 mg or $\frac{1}{24}$ $\frac{1}{12}$ gr /dose) and the patient observed for several days after each increment before going on to the next level Caution should be exercised in older patients to avoid lowering the pressure too rapidly this is true also of those patients with evidence of atheroma in the cerebral or coronary arteries since acute hypotension may result in thrombosis of these vessels
- b Pentolinum Tartrate N N D (Anslysen[®]) The initial dose is 1 2 mg ($\frac{1}{60}$ $\frac{1}{30}$ gr) of the salt given subcutaneously If no untoward effect occurs the dose can be gradually increased beginning on the second day by increments of 0 5 mg ($\frac{1}{120}$ gr) On discharge from the hospital the average patient receives 5 to 10 mg ($\frac{1}{12}$ $\frac{1}{6}$ g) per day in divided doses The same precautions noted for hexamethonium are to be observed

Following discharge the patient should be seen at frequent intervals and the dose adjusted so as to achieve the desired effect without undue faintness or side effects In some patients in order to prevent a postural hypotension which may produce severe symptoms during this period it may be necessary to have the patient lie down for one hour after each injection In many of these patients however tolerance gradually develops although marked hypotension may still occur on standing the patient may be able to sit or walk immediately after an injection Patients should be warned to avoid motionless standing for an hour or so after an injection avoiding waiting in line for a bus and similar activities should be particularly condemned

- 4 Acute hypertensive emergencies Give hexamethonium intravenously at a rate of approximately 1 mg ($\frac{1}{60}$ gr) per minute to a total dose of 10 to 20 mg ($\frac{1}{6}$ $\frac{1}{3}$ gr) depending upon the response of the patient to these doses 2 5 mg every 1 hour The most important of the emergency situations treated in this way is a pulmonary edema associated with a marked rise in blood pressure occurring in hypertensive patients with left ventricular failure Pulmonary edema may improve dramatically under this circumstance Great caution must be used to give the drug slowly and to discontinue administration when a moderate fall in pressure has been achieved A further fall in blood pressure may occur for a time after the drug is withdrawn
- 5 Side effects and hazards of ganglionic blocking agents

ANGINAL SYNDROME (Angina Pectoris) (code No 401)

The diagnosis of angina pectoris must be based upon positive diagnostic criteria not available by exclusion. The diagnosis depends upon proper interpretation of a careful history and a accurate evaluation of the credibility of the patient.

Diagnosis

The cardinal symptom of the anginal syndrome is pain induced by anything that increases the work of the heart (e.g. exertion, excitement, cold, heavy meals). Pain is usually substernal but may be precordial. The onset is sudden but not instantaneous. Its character is that of pressure or a squeezing sensation of significant but short duration, rarely lasting more than 15 or 20 minutes.

Treatment of the Acute Attack

A. Specific Measures (Nitrites)

1. Glyceryl Trinitrate U.S.P. B.P. (nitroglycerin) is the drug of choice. It acts in about 1-2 minutes. As soon as the attack begins place 0.3 mg ($\frac{1}{2}$ 200 gr.) tablet under the tongue and allow it to dissolve. The dose may be increased to 0.4-0.6 mg ($\frac{1}{2}$ 150 $\frac{1}{2}$ 100 gr.) if no relief is obtained from a small dose. Nitroglycerine may be administered whenever an attack occurs or may be administered to prevent an attack. It may cause headache and hypotension.
2. Amyl Nitrite U.S.P. B.P. 1 p.c. should be inhaled in about 10 seconds. This drug usually causes disagreeable reactions of flushing of the face, pounding of the pulse and sometimes dizziness and headache. The reaction may be minimized by inhaling the drug from a distended bag by partially pinching the crushed pea before the nose. The patient should learn how to vary the amount of drug he wishes to inhale.
3. Longeracting nitrite and other drugs have no place in the therapy of the acute attack.
4. Alcohol 30-60 cc (1-2 z) of whiskey, brandy, etc. may be a helpful home remedy.

B. General Measures. Rest is the most important therapy in an attack. The patient should cease any exertion and should stand still until he lies down. Once he detaches himself of pain and until the attack is over. This generally is the natural reaction of most patients but some try to work through the attack. Patients should be warned against this.

Prevention of Further Attacks

A. Specific Measures

1. Drug

Longeracting nitrite P. t. e. ythritol Tetranitrate N.N.D. (P. rit. te²) must be the most effective of the three. The suggested dose is 10 mg ($\frac{1}{8}$ g.) tid.

- a. Glyceryl Trinitrate U.S.P. B.P. (nitroglycerine) 0.3-0.6 mg ($\frac{1}{2}$ 200 $\frac{1}{2}$ 100 gr.) under the tongue just before activity.

Xanthin This drug may be of some benefit only in the aged (see page 204).

aggravate the degree of existing hypertension and increase the load on the heart and kidney. Attempts have been made to treat hypertensive patients with psychoanalytic methods but the effect on the blood pressure has been poor even though symptoms may often be improved. Reversal of hypertension following psychotherapy has been extremely rare. Attention to the emotional needs of the patient is an important adjunct to other methods of treatment but should not be the sole method of treatment except in the mild benign forms of the disease in which drug or surgical therapy is not indicated.

Other Method of Treatment

A Sedation Nervous tension is frequently found in the hypertensive patient and may aggravate his illness. In many cases sedation either used alone or as an adjunct to other forms of medical therapy will be of decided benefit. Phenobarbital is the drug most commonly used. Dosage 15-30 mg ($\frac{1}{4}$ to $\frac{1}{2}$ gr) t.i.d. to q.i.d.

B Drugs which have evoked little general enthusiasm despite occasional favorable results because of the unpredictable effects on the hypertension and the high incidence of unpleasant side effects include Dibenamine[®], the dihydrogenated ergot preparations, Toluoline Hydrochloride U.S.P. (Priscoline[®]) and potassium thiocyanate and the long acting nitrites.

Treatment of Complications

The cardiac, cerebral and renal complications of hypertension are discussed under congestive failure (see page 181), angina pectoris and myocardial infarction (see page 163), cerebral hemorrhage and thromboses (see page 349) and renal failure (see page 301).

Headache

The headache of hypertension is largely on an emotional basis. Suggestion and explanation are often helpful. Hypotensive drugs are most effective in relieving even the headache associated with the malignant or pre-malignant phases of hypertension.

CORONARY HEART DISEASE

Coronary Insufficiency

Coronary insufficiency is a dynamic concept which is connected with the balance between the blood flow in the coronary arteries and the demands of the myocardium for blood. It exists whenever the requirement of the myocardium for oxygenated blood exceeds the flow of blood of the myocardium at any instant.

Coronary insufficiency may be acute and transient in which case it is called a *giant's infarction* or it may be subacute moderately protracted and without myocardial necrosis but clinically recognizable. The latter form has been called coronary failure by Blumgart and his associates and is thought to represent occlusion of a relatively minor coronary artery with sufficient collateral circulation to prevent myocardial necrosis. Treatment is similar to that of a mild infarction.

- 2 Abdominal support Obese patients with protuberant abdomens who have angina may have fewer attacks following the use of proper abdominal support the mechanism is not clear. The Kerr Lagen b It is designed for this purpose.
- 3 Surgical procedures These have been employed only in patients with severe incapacitating angina pectoris in whom medical treatment has failed the results to date have been inconclusive. Studies now under way on the ligation of the internal mammary arteries have yet to be evaluated.
- 4 Production of myxedema by means of thiouracil compounds or radio active iodine (I^{131}) (see p. 372). The objective is to reduce the work of the heart. Good results have been reported in about half of the cases of intractable angina but this method should not be used until prolonged rest and attention to the emotional needs of the patient have ruled out a transient reversible coronary insufficiency.

B General Measures

- 1 The patient must avoid all habits and activities that he knows will bring on an attack.
- 2 Treatment of existing disorders especially anemia which may lead to increased cardiac ischemia.
- 3 Rest Most patients with angina do not require prolonged bed rest but rest and relaxation are beneficial. Adequate mental rest is also important.
- 4 Diet Obese patients should be placed on a reducing low animal fat diet and their weight brought to normal or slightly subnormal levels.
- 5 Tobacco is best avoided or used in moderation because it produces tachycardia and elevation in blood pressure.
- 6 Hypercholesterolemia has been shown to accelerate atherosclerosis in man and to be essential in its production in animals. If the serum cholesterol exceeds 260 mg % in a patient with angina pectoris an attempt should be made to lower it by diet with total calories containing about 25% fat (50% vegetable and 40% animal). If this is unsuccessful beta sitosterol (Cytellin®) may be added 1 or 2 Tbsp immediately before each meal. It has not been shown however that lowering the serum cholesterol level will reverse the atherosclerotic process.

ACUTE MYOCARDIAL INFARCTION (code No. 430.516.7)

Myocardial infarction is due to necrosis of a portion of the cardiac muscle as a result of impairment of its blood supply. This impairment usually results from occlusion or thrombosis of a coronary artery but it may result from impaired blood flow as a result of shock or acute anemia from any cause. Myocardial infarction varies quantitatively from histologically to massive necrosis. The infarction may be essentially asymptomatic.

The onset of an angina pectoris may be associated with coronary occlusion even though infarction does not occur (if the collateral blood flow is adequate). The prognosis is better than previously thought.

failure of the digitalis with are (see page 197)

- 3 Stokes-Adams attack with heart block is an emergency (see page 180)
- 4 Thrombo-embolism phenomena are common during the course of myocardial infarction. If anticoagulants have not been given they should be promptly administered (see page 215)
- 5 Extension of the infarction. When a patient has a big infarction of the myocardium, it should be suspected and confirmed. It ought to be treated in the hospital and in other clinical facilities. The same method of treatment applies to the original infarction but the prognosis is equal.

Activity Status in Congestive

The minimum period of rest should be at least 3 weeks after the attack has been over. The patient should be encouraged to approach gradually 6 weeks. The program for most patients is 1 month of complete rest, 1 month of low activity, and a third month of rest. It is difficult to put this into words. The amount of rest should be individualized according to the severity of the myocardial infarction and the response of the patient.

The patient should be permitted to walk freely about the room for about 7-10 days after the first attack. On the 11th day, the patient should be permitted to walk for 15 minutes. He should remain in the armchair with gradually increasing periods of walking slowly and with increasing heart rate. The patient should be able to walk for 15 minutes. The patient should be able to walk for 15 minutes. The patient should be able to walk for 15 minutes.

CHRONIC RHEUMATIC HEART DISEASE

Rheumatic heart disease is one phase of the rheumatic fever cycle. The stage of symptomatic valvular heart disease with the diagnosis is the late period between the subacute and acute rheumatic fever and the terminal phase of cardiac failure. The physiological decompensation of the late phase is much more possible.

Management of Asymptomatic Valvular Heart Disease

A Prophylaxis

- 1 Prevention of infection of the heart by
 - a Antidysrhythmic to prevent arrhythmias
 - b Continuous antibiotic prophylaxis in selected cases
 - c Prompt and adequate treatment of hemolytic infection
- 2 Prophylactic measures against the development of
 - a Edema
 - b High blood pressure
 - c Prevention of bacterial infection

B General Measures

- 1 Prevention of angina to anticipate a later period when the patient is more severely and significantly limited
- 2 Early recognition of disturbance of the old function and the presence of arrhythmias and the use of appropriate therapy
- 3 Maintenance of general health with good habits and diet
- 4 A plan of healthy diet and physical rest

is delayed and appears after the pain has subsided

a Vasopressor drugs Present evidence suggests that vasopressor drugs (sympathetic amines) may elevate the blood pressure and decrease mortality in myocardial infarction associated with shock. Shock must be treated early to achieve the best result. For details of the use of vasopressor drugs see page 30.

b Digitalis A hypotonic myocardium often accompanies acute myocardial infarction and shock may be associated with an increased venous pressure. Some investigators now favor digitalization in the shock of acute myocardial infarction. Digitalization can be accomplished in congestive heart failure. The increased cardiac output increases coronary flow and the peripheral myri-

c Treatment of cardiac arrhythmias Shock may be the result of undetected ectopic tachycardia or other arrhythmia and prompt treatment of the arrhythmia (see below) and cardiac arrhythmias may be lifesaving.

d Iron and ascorbic acid transfusions These have not been very effective but should be kept in mind as adjuncts.

6 Anticoagulant therapy This is a controversial matter in the milder as well as in the severe cases of myocardial infarction. In severe cases of myocardial infarction anticoagulants are generally recommended. For technique see page 215.

7 Sedation Adequate sleep is essential in patients with myocardial infarction. Sedation should be used as necessary to provide sufficient sleep and morphine derivatives should not be withheld in the first few days if the patient indicates.

B Follow-up Careful clinical observation is mandatory to determine the extent of the infarction and the appearance of complications or symptoms requiring treatment.

C Treatment of Complications

1 Cardiac failure If cardiac failure develops, treatment should be initiated immediately. Oxygen, low sodium intake, mercurial diuretics, and digitalis are the essential treatments. The patient should be digitalized in such a manner as to avoid toxic reaction. If possible, rapid digitalization should be avoided unless the failure is urgent. If the cardiac failure is mild and manifested solely by pulmonary congestion and increased dyspnea, restriction of sodium and the administration of mercurial diuretics may be sufficient. Digitalis is avoided by some authorities because of the hazard of ventricular arrhythmias but its well-controlled administration should not be deferred if cardiac failure is demonstrated.

2 Arrhythmias

a Ventricular premature beats These are common and increased mortality of the delayed myocardial infarction may predispose to ventricular tachycardia. Quinidine sulfate is the drug of choice (see page 200). An alternative to quinidine is procainamide (see page 205).

b Ventricular tachycardia is a emergency (see page 178).

c Atrial fibrillation is usually transient. If this persists, if the patient tolerates it poorly or if congestive heart failure

diag. is of mitral stenosis is difficult under these circumstances and the surgical mortality high.

AORTIC STENOSIS (code No 499)

Operat e reh f fm han la t e t o s ha be n s
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Surgical Treatment

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BACTERIAL ENDOCARDITIS

(Suba ute code No 450-100 0) (A ute code No 450-100)

These are infections of our gastrointestinal tract of the
 the (most frequently the valve) regurgitation associated with
 how rapidly the small intestine moves. The most common
 of the gastrointestinal tract is the St. Louis epidemic
 of the which the intestinal bacteria enter the
 bacteria which may cause many infections of the
 pneumonia may cause bacterial endocarditis due to
 the staphylococci, beta hemolytic streptococci, Hemophilus
 influenzae and gonococci.

D. ENCE

Symptoms are all those of peritonitis signs include pallor petechial hemorrhages pleural effusions of lung and fluid of uli congenital heart disease Do not make diagnosis of subacute bacterial endocarditis in the absence of valvular congenital heart disease without repeated positive blood cultures showing the same organisms Do not exclude the diagnosis of bacterial endocarditis with repeated negative blood cultures in low cultures Occasional typical clinical cases will be found to have peritendinitis gutti blood culture Under the hood the symptoms are static and empirically the patient is damaged to heart valve

MITRAL VALVULAR DISEASE

This is the most common of valvular lesions. It takes from 3 to 5 years for mitral stenosis to develop. mitral insufficiency may occur alone or more commonly in combination with mitral stenosis.

MITRAL STENOSIS (code No. 498)

In view of the excellent results obtained following mitral valvulotomy the signs of mitral stenosis should be clearly appreciated.

Diagnosis

A Signs of Uncomplicated Mitral Stenosis The most important of these are (1) a mid diastolic long murmur always associated with presystolic accentuation if there is sinus bradycardia and usually associated with a thrill (2) a systolic flutter and an opening snap and (3) an apical systolic murmur at the apex or one which is short and Grade II or less.

If pulmonary hypertension is present its signs and those of associated right ventricular hypertrophy may be demonstrated.

B Exclusion of Mitral Insufficiency Mitral incompetence must be excluded if possible. The mitral valve is operable only if the patient's condition is due to a mechanical obstruction of the mitral valve. If there is no systolic murmur in the presence of the signs of mitral stenosis mitral incompetence is exceedingly unlikely. If there is a loud pan systolic murmur at the apex in association with an accentuated often early 3rd heart sound a soft 1st sound and no opening snap the diagnosis of predominant mitral incompetence is likely even if a short mid diastolic murmur can be heard at the apex. Left ventricular hypertrophy in the ECG should make every physician in recommending surgery for mitral stenosis because of the likelihood of significant mitral incompetence unless hypertension or an aortic valvular lesion is present. If there is a moderate systolic murmur at the apex the diagnosis must rest on a consideration of the total findings.

Surgical Treatment

The course of mitral stenosis is highly variable and in view of the mortality of mitral valvulotomy (3.5%) surgery is not advised in mild cases with slight exertional dyspnea and fatigue only. Indications for surgery include the following:

- 1 Signs of mitral stenosis with a pliable valve (opening snap snapping 1st sound)
- 2 Uncontrollable pulmonary edema
- 3 Disabling dyspnea and occasional pulmonary edema
- 4 Evidence of active pulmonary hypertension with right ventricular hypertrophy and early congestive failure
- 5 Systemic and pulmonary emboli
- 6 Increased pulmonary arteriolar resistance with marked dyspnea and increased P_2 . The patients are apt to develop right heart failure and emboli
- 7 Right heart failure with atrial fibrillation tripping in incompetence when secondary to marked mitral stenosis. The

Approximate Dosage Schedules

Penicillin Inhibition (Bacterialidal at 72 Hrs) (Unit per cc)	Total Penicillin Dose per 24 Hrs (Millions of Unit)
< 0.1	1.2 (penicillin procaine)
0.1-0.5	3.4 (aqueous)
0.5-0.9	4.5 (aqueous)
1.0-5.0	6.20 (aqueous)
> 5.0	20.500 (aqueous)

When bacteremia deferves peristaltic dose should be doubled and redoubled until full response occurs. Alternatively synergistic treatment with 2.0 mg of tetracycline may be used. When high concentration of penicillin is required, Penicillin NND (B. mid[®]) 0.5 Gm (7½ gr) every 6 hours may be used to inhibit renal excretion.

- Streptomycin Sodium U.S.P. Intramuscular Injection is the method of choice and gives a good level as those obtained by intravenous injections. Large doses are advised: 0.5-1.0 Gm dissolved in 4 cc (1 cc) distilled water + 1-2% procaine in 1 M. Every 6 hours should be given. Observe for toxicity.
 - Combination of penicillin and streptomycin. Preliminary evidence suggests that penicillin (5 million units/day) + tetracycline (2 Gm/day) may be the optimal treatment for infections due to Streptococcus faecalis and all of the heart (2 weeks). The treatment of endocarditis due to staphylococci of Streptococcus viridans.
 - Chlortetracycline Hydrochloride U.S.P. (Auromycin[®]) Oxytetracycline U.S.P. (Terramycin[®]) Tetracycline U.S.P. (Achromycin[®]) Chloramphenicol U.S.P. (Chloromycetin[®]) and Erythromycin U.S.P. (Erythrocin[®]). While these drugs may suppress the progress of subacute bacterial endocarditis their use is frequently followed by relapse. Whenever possible drugs exhibiting more pronounced bactericidal activity e.g. penicillin and streptomycin should be the first choice in treatment. The exact dosage and effectiveness of therapy have not been established. Nausea and vomiting result frequently from the oral administration of chlortetracycline and may interfere with treatment. In such cases the drug must be given intravenously in doses of 50-100 mg or more every 6 hours.
- Although streptomycin is clinically generally inhibited by chlortetracycline oxytetracycline and tetracycline treatment with these drugs of endocarditis due to staphylococci is generally ineffective.
- Other drugs: Neomycin Sulfate U.S.P. (Myfrin[®]) Bacitracin U.S.P. and Polymyxin B Sulfate U.S.P. (Aerosporin[®]) may be used alone in combination with other drugs where the organism is insensitive to the toxic antibiotics (see page 514).

Treatment

A. Specific Measures The most important consideration in the treatment of bacterial endocarditis is a bactericidal concentration of one or more antibiotics in contact with the infecting organisms which are often localized in avascular relatively inaccessible foci. Penicillin because of its high degree of bactericidal activity against the great majority of bacteria which produce bacterial endocarditis and because of its low incidence of side reactions is by far the most useful drug. Synergistic combinations of penicillin with other antibiotics have often proved valuable. Few cases have been cured by bacteriostatic drugs such as chlortetracycline (Aureomycin®), oxytetracycline (Terramycin®), tetracycline (Achromycin®), chloramphenicol (Chloromycetin®) and erythromycin (Erythrocin®) used alone. Positive blood cultures are invaluable to confirm the diagnosis and to guide treatment and should be combined with tests of sensitivity of the infecting organism to various antibiotics or combinations of antibiotics. Hence one or more blood cultures should be obtained daily for 3 to 5 days before instituting treatment except in desperately ill patients or patients with acute bacterial endocarditis. To avoid further heart damage treatment should not be further delayed.

1. **Penicillin** This drug must be given parenterally in bacterial endocarditis in order to gain effect. The dose of penicillin used depends on the sensitivity of the organism and this is determined by doing in vitro sensitivity tests. About 90% of strains of *Streptococcus viridans* from cases of subacute bacterial endocarditis have been found to be inhibited in vitro by 0.1 unit of penicillin per cc or less. However some are quite resistant requiring 5 to 10 units or more.

A minimum serum concentration many times greater than the apparent in vitro sensitivity of the organism must be produced to insure a bactericidal concentration in the circulation. In patients in whom positive blood cultures are not obtained or when sensitivity tests are not available 5 to 10 million units of penicillin should be given daily. There are three alternative methods of administration.

- a. **Procaine penicillin** For organisms sensitive to less than 0.1 U per ml of penicillin give 500,000 to 1,000,000 units of penicillin procaine I.M. twice daily.
- b. **Intermittent administration** For organisms sensitive to 0.1 U per ml of penicillin or more intermittent intramuscular injections of aqueous penicillin solution every 3 to 6 hours.
- or c. **Continuous parenteral administration** If the total daily dose is approximately 5 million or more units of penicillin per day administration usually best accomplished by a continuous intramuscular drip (occasionally intravenous drip). The antibiotic can be dissolved in 1000 to 2000 cc of physiological saline solution or glucose solution.

Approximate Dosage Schedules

Penicillin Inhibition (Bactericidal at 72 Hr) (Unit per cc)	Total Penicillin Dosage per 24 Hr (Millio of Units)
< 0.1	1.2 (penicillin procaine)
0.1 to 0.5	3.4 (aqueous)
0.5 to 0.9	4.5 (aqueous)
1.0 to 5.0	6.20 (aqueous)
> 5.0	20.500 (aqueous)

When bacteremia is definitely present dosage should be doubled and continued until favorable response. Alternatively synergistic treatment with tetracycline or chloramphenicol may be used. When high concentration of penicillin is required Procaine Penicillin (B.N.D.®) 0.5 Gm (7½/2 g) every 6 hours may be used to inhibit infection.

Streptomycin Sulfate U.S.P. Intramuscular Injection is the method of choice and gives a good response. The dose is 1.0 Gm daily by I.V. injection. Large doses are advised 0.5 to 1.0 Gm daily in 4 cc (1 dr) distilled water + 1 cc 2% procaine I.M. every 6 hours should be given. Observe for toxicity.

3. Combined penicillin and streptomycin. Penicillin G (5 million unit/day) + streptomycin (2 Gm/day) may be the optimal treatment for infective endocarditis due to Streptococcus viridans and also for the short (2 weeks) treatment of endocarditis due to sensitive strains of Streptococcus viridans.
4. Chlorotetracycline Hydrochloride U.S.P. (Achromycin®), Oxytetracycline U.S.P. (Terramycin®), Tetracycline U.S.P. (Achromycin®), Chloramphenicol U.S.P. (Chloromycetin®) and Erythromycin U.S.P. (Erythrocin®). While these drugs may suppress the progress of subacute bacterial endocarditis, their use is frequently followed by relapse. When ever possible drugs exhibiting marked pronounced bactericidal activity such as penicillin and streptomycin, should be the first choice in treatment. The exact dosages and effectiveness of the drugs have not been established. Naïve and untreated patients frequently form the oral administration of chlorotetracycline and may interfere with treatment. In such cases the drug must be given I.V. in doses of 50-100 mg or more every 6 hours.

Although Streptococcus faecalis is generally inhibited by chlorotetracycline, oxytetracycline and tetracycline, treatment with these drugs of endocarditis due to this organism is generally ineffective.

5. Other drugs. Nafcillin Sulfate U.S.P. (Mycifradin®), Bacitracin U.S.P. and Polymyxin B Sulfate U.S.P. (Aerospirin®) may be used also in combination with other drugs when the organism is sensitive. It is toxic and but (see page 314).

- 6 Combined therapy In infections due to highly resistant organisms synergistic pairs of antibiotics as determined by tests of bactericidal activity in the laboratory may be used (see page 496) Combined therapy should never be attempted without adequate laboratory control
 - 7 Duration of treatment The suggested duration of therapy by various authors is 2-8 weeks Most patients should be treated for 3-4 weeks after sterilization of the blood stream After therapy has been discontinued the patient should be carefully observed for recurrence by taking repeat blood cultures
 - 8 Recurrences Most recurrences are observed within a week or two of the end of therapy Occasional cases relapse months later The diagnosis of recurrence must not be made on the return of fever and embolic phenomena alone these may occur for up to 6-8 weeks after therapy has ceased Positive blood cultures are essential for the diagnosis of recurrence Before re-treating again determine the sensitivity of the organism and then give treatment with higher dosages for a longer period of time or use a different antibiotic About 70-75% recurrences are now being reported
 - 9 Anticoagulants It is generally agreed that the use of heparin or Dihydroxycoumarin U.S.P. (Dicumarol[®]) in the treatment of subacute bacterial endocarditis is unnecessary and may be dangerous
 - 10 General Measures General supportive measures as for any severe infection with fever should be given
- C Complications and Treatment

- 1 Infarction Caused by emboli breaking off from the infected areas The infarctions usually occur in organs in the systemic circulation but if the endocardial lesion is on the right side of the heart the embolus may be to the pulmonary circulation Treatment is symptomatic
- 2 Cardiac failure (uncommon) Acute myocarditis or swelling of the heart valves may precipitate congestive failure When giving large quantities of penicillin as sodium salt one may give significant amounts of sodium ion Therefore when treating a case of subacute bacterial endocarditis with congestive failure or possible failure use calcium or potassium penicillin (See congestive failure page 181)
- 3 Anemia The anemia if severe should be treated by whole blood transfusions (see page 247)
- 4 Uremia May result from focal embolic nephritis or glomerulonephritis (see page 293)

Prophylaxis

A high percentage of cases of endocarditis arise after dental procedures or surgery of the oropharynx and genitourinary tract Therefore all patients with valvular or congenital heart disease who are to have any of these procedures should be given penicillin prophylactically A satisfactory schedule is as follows: Penicillin G U.S.P. 1,000,000 units daily for 2 days before procedure on the day of the procedure and for 2 days after the procedure

CARDIAC ARRHYTHMIAS*

Cardiac arrhythmias are common in every physical practitioner and a thorough knowledge of their diagnosis and management is essential. Clinical manifestations vary from trivial palpitation to a clinical state of the utmost urgency as when ventricular tachycardia complicates acute myocardial infarction.

Relation of Symptoms

The symptoms produced by an arrhythmia depend upon the underlying state of the heart and the duration of the arrhythmia. Even a normal heart may fall if the ventricular rate is rapid enough if the arrhythmia lasts long enough. Tachycardia which may be well tolerated by an individual may produce severe pulmonary edema in another (e.g. a patient with tight mitral stenosis). The physical manifestations develop when the tachycardia is sustained or when the tachycardia is associated with a severe effort to treat the arrhythmia.

DISTURBANCES OF ATRIAL ORIGIN

ATRIAL PAROXYSMAL TACHYCARDIA (Code No. 422)

In the diagnosis of arrhythmia it is thought that an ectopic focus within the atrium is the cause as the pattern of the heart and discharge impulses at the atrial focus from 120 to 320 per minute usually between 170 and 210 per minute. The rhythm is absolutely regular and is affected by inspiration movement or circadian pressure (unless this abolishes the attack). Atrial tachycardia usually begins suddenly unless it implicates even a predisposition. At least half of the cases occur in individuals with organic heart disease. During an attack is rare but characteristic of the arrhythmia. The rhythm may occur when the rhythm produces last of a period of day. The attacks are often produced by emotional tension, a common phenomenon in individuals and at times are related to a circadian rhythm. The A-V node of the atrium is the Wiggers-Parkinson-White syndrome. R current attacks are frequent so that the problem of preventing attacks is an important aspect of the treatment of the individual attack.

Treatment of the Acute Attack

The basic of heart disease is sufficient to cause a myocardial infarction subspontaneously and the physical should not underestimate the importance of the disease. Patients should be made to stop the attack quickly if it persists for several days if a definite syncope or anginal pain develop or if the patient is dying a cardiac death.

A Manual of Methods. A variety of methods have been used to interrupt attacks and the patient may learn to do this himself. These include the Valsalva maneuver (holding breath and increasing intra-abdominal pressure) stretching of the arms and body lowering the head between the knees and holding.

B Vagal Stimulation

- 1 **Carotid sinus pressure** With the patient relaxed in the semi recumbent position firm but gentle pressure and massage should be used first over one carotid sinus for 10 to 20 seconds and then over the other. Pressure should not be exerted on both carotid sinuses at the same time. Continuous auscultation of the heart should be carried out so that pressure is stopped as soon as the attack ceases. Carotid sinus pressure will interrupt about half of the attacks especially if the patient has been digitalized.
- 2 **Bilateral eyeball pressure** has been recommended but it is rarely as effective as carotid sinus pressure and carries the risk of producing a detached retina.
- 3 **Induced vomiting** (except in cases of syncope, anginal pain or severe cardiac disease)

C Drug Therapy If mechanical measures fail and the attack continues (particularly if the above symptoms are present) drugs should be employed. There is no unanimity of opinion about the most effective drugs but the following are satisfactory.

- 1 **Quinidine Sulfate** U S P B P (see page 200)
- 2 **Neostigmine** U S P (Prostigmin®) 1 mg subcutaneously
- 3 **Digitalis** orally or if no digitalis has been given in the preceding 2 weeks intravenously
- 4 **Procainamide Hydrochloride** U S P (Pronestyl®) (see page 205) Continuous electrocardiograms or continuous monitoring of the heart rate and blood pressure is essential.
- 5 **Methacholine Chloride** U S P B P (Mechoyl®) 10 mg subcutaneous is often effective but produces very unpleasant side effects and should rarely be used.
- 6 **Syrup of Ipecac** U S P 4 to 8 cc may be used to induce vomiting. It may be repeated if unsuccessful.

Prevention of Attacks

- A** Attempt to find and remove the cause especially emotional stress, undue fatigue or excessive use of alcohol or tobacco.

B Drugs

- 1 **Quinidine Sulfate** U S P B P 0.2 to 0.8 Gm (3 to 9 gr) t i d to q i d may be used to prevent frequent and troublesome attacks. Begin with small doses and increase if the attacks are not prevented and toxic effects do not occur.
- 2 Should quinidine fail or if it is not tolerated full digitalization followed by digitalis in maintenance doses may prevent or decrease the frequency of attacks (see page 197).
- 3 **Procainamide Hydrochloride** U S P (Pronestyl®) in a maintenance dosage of 250 to 500 mg t i d may be tried if the above methods are not successful.

NODAL PAROXYSMAL TACHYCARDIA (code No. 422)

This resembles atrial tachycardia except that the ectopic focus is in the A-V nodal tissue. Although the electrocardiographic or clinical distinction between atrial and nodal paroxysmal tachycardia is not possible in which case the term supraventricular tachycardia is used. Treatment is carried out along the same lines as for atrial tachycardia (see page 173).

ATRIAL FLUTTER

(Paroxysmal code No 423) (Chronic code No 424)

This a rhythm is due to impulses which rise from an irritability of the atrial muscle at rates of 250 to 350 per minute. The ventricular rate is usually one half the atrial rate (2:1 conduction) but it may be 1:1 (3:1 or 4:1 conduction) or very rarely a 1:1 conduction may occur with a very rapid ventricular rate. The ventricular rate is usually regular but if there is a significant A-V block it may be irregular and may imitate atrial fibrillation. Atrial flutter usually results from any of the common causes of heart disease and it may occur infrequently in the absence of heart disease. It may be produced by quinidine during the treatment of atrial fibrillation.

Treatment

A Treatment of Paroxysmal Flutter Similar to the treatment of paroxysmal tachycardia except that digitalis and quinidine are the drugs of choice. The arrhythmia tends to become established more often than do the other cardiac tachycardias. Prophylaxis of recurrence attacks is arranged similarly to that described for atrial tachycardia (see page 173).

B Treatment of Chronic Atrial Flutter

- 1 Digitalis is the drug of choice. It increases the A-V block and prevents a 2:1 or 1:1 conduction. Usually half of the usual atrial fibrillation sinus rhythm results from full digitalization. If atrial fibrillation remains fixed it has been produced by digitalis. Quinidine sulfate may be added to convert to sinus rhythm. Digitalis may be given by any one of the methods (see page 197). Other medication is usually sufficient although the intravenous route may be used if the situation is critical. Digitalis must often be given in larger doses than is usually required for cardiac failure. When a fixed 4:1 conduction is produced by digitalis a slightly increased dose may convert the flutter to atrial fibrillation or sinus rhythm.
- 2 Quinidine Sulfate U.S.P. B.P. This drug should not as a rule be used to treat atrial flutter unless the patient is fully digitalized with a slow ventricular rate because of the danger of producing a 1:1 conduction. If digitalis resists in only a 4:1 conduction or produces atrial fibrillation which does not respond satisfactorily to sinus rhythm quinidine may be given (see atrial fibrillation below).

ATRIAL FIBRILLATION

(Paroxysmal code No 425) (Chronic code No 426)

A common arrhythmia due to ectopic impulse arising from the atrium at a very rapid rate (400-500) they are not followable at all pathways. The ventricular rate is always irregular in character usually varying between 110 and 160 but may be slow or faster depending upon the degree of A-V block. The chamber for measurement is usually the first, usually a so-called with a cardiac catheter. Usually the small mitral valve diastolic pressure and hypotension are noted and thyrotoxicosis. The paroxysmal type may

occur without apparent reason in normal individuals in apparently normal hearts during acute infectious diseases following surgical operations especially of the lungs and particularly in thyrotoxicosis

Treatment

A Treatment of Paroxysmal Atrial Fibrillation

1 Specific treatment

- a Digitalis is the drug of choice in paroxysmal atrial fibrillation especially when this arrhythmia occurs in individuals with organic heart disease (particularly mitral stenosis) with rapid ventricular rates or when the symptoms or signs of cardiac failure have appeared. If there is doubt as to whether one should use quinidine or digitalis first digitalis should be given. This is because it controls the ventricular rate by producing an A-V block which is the immediate objective of treatment in such a case. The objective of treatment with quinidine is to abolish the atrial ectopic rhythm and it is quite safe to wait until the ventricular rate is brought under control with digitalis. Give full digitalizing doses (see page 197) with the objective of slowing the ventricular rate to 70 to 80 per minute and avoiding toxic manifestations. In paroxysmal fibrillation there is no clear evidence that the use of digitalis will result in established fibrillation.
- b In those cases where an attack of atrial fibrillation persists in an otherwise normal heart with a ventricular rate under 180 and with no other symptoms or signs of cardiac failure quinidine sulfate may be used at once to convert the rhythm to sinus rhythm.

If the ventricular rate becomes very rapid or if symptoms of dyspnea, anginal pain or a very rapid pulse are produced conversion with quinidine should be temporarily suspended and digitalis given.

- 2 Prophylaxis of paroxysmal fibrillation. The principles and procedure are the same as for atrial paroxysmal tachycardia (see page 173).

B Treatment of Chronic Atrial Fibrillation

1 Drugs

- a Digitalis. The long digitalization is the first step (see page 197). The patient is then usually placed on maintenance digitalis indefinitely. The object of digitalization is to slow the ventricular rate and to improve myocardial efficiency.
- b Quinidine Sulfate U.S.P. B.P. Quinidine is used to abolish the ectopic rhythm once the ventricular rate is controlled with digitalis. It potentially has a hard use and should be used only in carefully selected cases by a physician thoroughly familiar with the drug and by a method which ensures close medical supervision (preferably in the hospital) while conversion to sinus rhythm is being attempted. CAUTION See page 200 for dangers of quinidine.

- 2 Conversion of chronic atrial fibrillation. Opinion is divided but the following indications for conversion of atrial fibrillation serve as a general guide. Each case must be individually evaluated. In general conversion is attempted when either it is

thought that the patient will be better off with sinus rhythm than with fibrillation.

- a Atrial fibrillation persisting after thyrotoxicosis has been treated surgically or by other means.
- b Atrial fibrillation of a few weeks' duration in an individual with or only light thyroid disease.
- c Atrial fibrillation associated with frequent embolophthalmia.
- d Refractory case of flutter induced by the treatment of fibrillation. Severe palpitations due to inability to decrease the ventricular rate with digitalis therapy may be obvious only on exercise.
- f Atrial fibrillation appearing for the first time postoperatively in patients with a technically successful mitral valvulotomy.

DISTURBANCES OF VENTRICULAR ORIGIN

VENTRICULAR PREMATURE BEATS (cod No 441)

A rhythmia in which top impulses are from any position in the ventricle to a premature beat. It is one of the most common arrhythmias and fits no individual with such a disturbance in which the occasional ectopic premature beat is good. Multiple ventricular premature beats or those arising from two or more different sites more often where they are in middle divided also they rise the question of lying on a y heart disease. Ventricular premature beats occurring in normal with coronary heart disease possibly when they produce symptoms or give place to few with the gates in the Thymopressage of the ventricular fibrillation. This spontaneously follow igmyo dial infusions. Ventricular premature beats may be associated with anxiety cases of alcoholism or fatigue. It is the most common rhythmia resulting from digitalis toxicity.

Treatment

A Specific Measures

1. Reassurance and good hygiene. If an associated disease is present and if the ectopic beats are of frequent and produce no palpitations no specific therapy indicated. Rest, sleep, and instruction of the patient concerning the relationship between his symptoms and his position are helpful.
2. Adjustment of digitalis dosage. If ventricular premature beats are due to digitalis toxicity stop the digitalis for 3-5 days or until the arrhythmia disappears and then resume medication in smaller dosage. At times however patients with cardiac failure who are receiving digitalis may develop ventricular premature beats which not due to digitalis toxicity but to increased digitalis at onset of cardiac failure. If it is due to the withdrawal of digitalis for 1 day a dose of the digitalis failure with other available methods.

(see page 182) In these circumstances the ventricular premature beats often disappear as the cardiac failure improves

- 3 Potassium salts 1-3 Gm (15-45 gr) q i d are often helpful in ventricular premature beats of digitalis origin.
- 4 Quinidine Sulfate U S P B P This drug should be used orally to abolish ventricular premature beats when they occur following acute myocardial infarction or when they occur in runs or from several foci in patients with heart disease

VENTRICULAR PAROXYSMAL TACHYCARDIA (code No 442)

A disorder in which impulse arise rapidly and fairly regularly in the ventricle at an average rate of 150-180 per minute. This is usually associated with severe myocardial damage especially myocardial infarction

Treatment

- A The Average Case
 - 1 Quinidine Sulfate U S P B P 0.4 Gm (6 gr) orally every 2 hours for 3 doses. If the attack is well tolerated and the patient is not in shock. If the attack continues and there is no toxicity from the quinidine increase dose to 0.6 Gm (9 gr) every 2 hours for 3 doses. This usually terminates the attack. If it does not give the drug I V or I M or change to procainamide.
 - 2 Procainamide Hydrochloride U S P (Pronestyl®) 0.5-1.5 Gm (7½-22½ gr) orally every 4 to 6 hours may be substituted for quinidine if the latter is ineffective or produces toxic symptoms.
- B The More Severe Case (or when other medication has failed)
 - 1 Quinidine Gluconate N F 0.8 Gm (12 gr) or 0.5 Gm (7½ gr) of quinidine base may be given I M and repeated every 2 hours for 2-3 doses.
 - 2 Procainamide Hydrochloride U S P (Pronestyl®) 0.5-1 Gm (7½-15 gr) may be given I M and repeated in 4 hours.
- C The Urgent Case
 - 1 Procainamide Hydrochloride U S P (Pronestyl®) 1 Gm (15 gr) given slowly I V at a rate not exceeding 100 mg (1½ gr) per minute. During the infusion continuous electrocardiograms and at least repeated blood pressure determinations are essential. Severe hypotension may result from this medication.
 - 2 Quinidine may be given I V as Quinidine Gluconate N F 0.8 Gm (12 gr) diluted with 50 cc of 5% glucose slowly (1 cc per minute) with continuous electrocardiograms and determination of blood pressure.
 - 3 Vasopressor drugs for shock. If shock is present as a result of ventricular tachycardia or results from the drug given I V it can be treated with vasopressor drugs as described under the treatment of shock (see page 30).
 - 4 Other drugs that have been described as occasionally helpful in ventricular tachycardia include
 - a Magnesium Sulfate U S P B P 10 cc of a 20% solution I V. Calcium salts should be readily available to counteract magnesium toxicity (see page

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VENTRICULAR FIBRILLATION (code N 445)

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SINO-ATRIAL BLOCK (cod No 416)

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Treatment

None is required generally. The causative factors should be eliminated if possible. The following drugs may be tried:

- 1 Atropine Sulfate U S P B P 0.6 mg (1/100 gr) q i d orally
- 2 Ephedrine Sulfate U S P Ephedrine Hydrochlorid B P 25 mg (3/8 gr) orally q i d

SINUS ARRHYTHMIA

Waxing and waning of the cardiac rate during the various phases of respiration. It is a normal phenomenon especially in childhood. Its main importance is its possible confusion with some important arrhythmias.

ATRIO-VENTRICULAR BLOCK

(Complete code No 436)

(Prolonged Conduction code No 434)

(Incomplete code No 435)

Delay in conduction through the A-V junctional tissues. The block may vary from mere prolongation of conduction time through progressive A-V block with varying degrees of block (2:1, 3:1, 4:1 conduction) to complete heart block (A-V dissociation). If complete heart block appears suddenly or if the ventricular rate slows abruptly in the presence of complete heart block, syncope may occur (Stokes-Adams syndrome).

Atrio-ventricular block occurs infrequently in normal hearts; it is usually due to organic heart disease. In young individuals it most commonly is due to rheumatism, fever, or diphtheria. In older patients coronary heart disease is a more common cause. Atrio-ventricular conduction defects may be caused by digitalis or quinidine therapy.

Treatment

In the absence of Stokes-Adams syndrome (see below) treatment of atrio-ventricular conduction defects is rarely accomplished except by elimination of drug if they are causative or by the subsidence of acute myocarditis. Prolongation of the atrio-ventricular conduction per se needs no treatment unless there is complete heart block with ventricular rate below 35/min. Cardiac failure or weakness may occur with slow ventricular rates. Ephedrine or Isoprenaline Hydrochloride U S P (Aludrin® is preferred) (see p. 181) should be given to increase the rate of the ventricular pacemaker.

STOKES-ADAMS SYNDROME (code N 455)

This syndrome refers to attacks of syncope as a result of ventricular standstill or more rarely ventricular tachycardia or fibrillation which occurs when an individual with no manifest atrio-ventricular conduction or one with a 2:1 atrio-ventricular block changes to a complete atrio-ventricular block. It is very serious.

nd then a d s dd death is common. If ca dia standstill is pro
l ng dfo m e th f w sec nds o v is o s may o cur. If the
h rt rate in complete hea t block ab ptly slows t under 20 an
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g) m y b g v n with a h d e f eph drine
- 2 I op oteranol Hydrochlo d U S P (Al dri e[®] Isup 1[®])
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q i d
- 3 Epinephrine Injection U S P If att ck ar fr que t and
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BUNDLE BRANCH BLOCK (code No 437)

(Right code No 4442 x37) (Left cod No 4443 x37)

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Testme t

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CONGESTIVE FAILURE

(Ca dia c Insufficiency code No 404)

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Diagno i

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increase in weight venous engorgement with increased venous pressure prolonged circulation time tender engorgement of the lower dependent edema and at times sacites

Congestive failure may be the end result of a wide variety of different types of heart disease and is therefore a functional diagnosis. The etiologic diagnosis must be made separately (e.g. rheumatic hypertension or coronary disease). Cardiac failure is often classified as acute or chronic and left sided or right sided. These divisions while often helpful are often arbitrary and may coexist.

Treatment

A Objectives of Treatment To increase the strength and efficiency of the myocardial contraction and to reduce the abnormal retention of sodium and water. The patient shares a significant responsibility in the management of his disease because treatment is long term and involves restrictions in diet and activity and because the patient's well being and productivity depend to a great extent on his willingness to cooperate.

B Correction of Causative Factors

1. Eliminate the causes of cardiac failure Cardiac failure may be reversible therefore in addition to carrying out the therapy to be outlined below specific search should be made for non cardiac causes of failure. These include thyrotoxicosis anemia myxedema nutritional disturbance (especially vitamin B deficiency) arteriovenous fistulae polycythemia vera and Paget's disease.

2. Eliminate precipitating factors Determine and eliminate if possible the factor precipitating the cardiac failure. Knowledge of the precipitating factor is important in treatment and in evaluating prognosis. The common factors precipitating failure include infection (especially respiratory) over exertion increased sodium intake discontinuation of medication (especially digitalis) onset of arrhythmia particularly with rapid ventricular rates (e.g. atrial fibrillation) myocardial infarction anemia and unknown factors.

C General Measures

1. Rest Rest in bed or sitting in a chair serves to decrease the work of the heart and to promote sodium diuresis. Morphine or barbiturate indicated as a welcome relief to a patient who has spent many sleepless dyspneic nights with his disease. Adequate rest should be maintained until compensation has occurred and then should be replaced by progressive ambulation. Most patients can sit at bedside comfortably with no more effort than is required for a bedpan.

Cardiac patients at bed rest are prone to develop phlebitis (See page 217 for prophylaxis). The duration of rest should be as long as necessary to permit the heart to regain reserve strength but should not be unduly prolonged so as to cause generalized debility of the patient.

2. Diet At the onset of therapy a few (4-6) small bland low caloric low residue meals with vitamin supplements are indicated. Restriction of sodium varies only in degree. Evaluation of the previous intake of sodium will provide a baseline upon which to gauge the degree of restriction required. Before drastic sodium restriction is instituted the

function holds but should be determined if the kidneys can conserve sodium (See page 187 for low sodium syndrome). In an occasional case 350 mg (5½ gr) or less of sodium may be the maximum tolerated without development of edema although sometimes the restriction is usually necessary only when failure is first treated. Vitamin supplements may be indicated. Restrictive diets and anoxia may lead to malnutrition and avitaminosis with a superimposed beriberi type of failure.

- 3 Digitalis (page 195) Digitalis increases the mechanical efficiency of the heart. In reabsorbed fluid output decreased, cardiac and ventricular distolipies and a fall in right atrial and peripheral venous pressure follow digitalis in patients with a diastolic failure. The glycosides available are qualitatively similar. They differ in speed of action, dosage, rate of excretion. It is advisable to be somewhat familiar with the parenteral and a rapid oral method. Rapid digitalization is indicated in atrial fibrillation with fast ventricular rate and in a fulminant pulmonary edema otherwise low digitalis tonics preferred (F method and dosage - see page 187).

- 4 Removal of sodium and water (See page 205)

a. Mercurial diuretics. Mercurial diuretic (CAUTION - see page 187-204) by decreasing the sodium and chloride reabsorption, the renal tubule. Clinical effect is noted in about 2 hours by the low molecular weight and is largely complete in 10-12 hours. Small quantities of mercurials (0.5 to 1 mg) may result in diuresis and should be started initially. They should be used in the morning so that their effect will have fully subsided by nightfall. Large doses may initiate massive diuresis with extensive fluid and electrolyte losses. This can be very distressing and can produce untoward symptoms particularly in the older age group. The toxicity of the mercurial diuretic is potentiated by giving Ammonium Chloride U.S.P. B.P. 2 Gm (30 gr) 4 times daily on the day before mercurial administration. The use of ammonium chloride for periods longer than 48 hours has no particular advantage and increases the danger of acidosis. The same is true of Acetazolamide N.N.D. (Diamox®) given 0.25 Gm twice daily for 2 or 3 days before the mercurial.

Merapton in Sodium N.N.D. (Thiomycin®) and phosphenamide Sodium N.N.D. (Merphidrin®) may be given subcutaneously and selected patients may be taught to inject themselves with the mercurial. The patient should be guided as to dosage by his weight gain. Close medical supervision is necessary to minimize the possibility of infection, fluid depletion, especially in the presence of renal and hepatic disease.

The mercurial diuretics may give very rapid results until the patient is dehydrated and diuresis has subsided and his dry weight is reached. Particular care should always be exercised to guard against a low sodium syndrome and hypokalemia (see page 15) which may occur.

particularly in patients who receive frequent mercurials while on a low sodium diet

- b Cation exchange resins [Carbacrylamine Resins N N D (Carbo Resin[®])] These substances are synthetic macromolecular compounds which undergo ionization and exchange ions for ions of similar charge. The acid potassium cation combination is probably best for cardiac failure. Resins are of greatest value in patients in whom urinary excretion of sodium is low and response to mercurials is poor. Eight Gm (120 gr) in a glass of water before and after each meal is the usual dose. The diet should contain 1.5-2 Gm (22½-30 gr) of sodium to achieve the most efficient use of resins and to prevent potassium depletion. Because the tendency to acidosis is increased, resins should be avoided in renal disease. The hazards of resin therapy are hyponatremia, hypokalemia, and acidosis. These hazards are minimized when the patient eats his full diet; if necessary, potassium salts can be given orally. The danger of calcium depletion during resin therapy is slight.
- c Paracentesis. Paracentesis of fluid in the chest and abdomen should be undertaken if respiration is embarrassed. Since sodium retention may occur as a result of fluid collections in the chest, abdomen, and legs, diuretics may occur following the procedure.
- d Mechanical measures. Venesection (in low output failure in the absence of anemia), Southey tubes, and acupuncture may be beneficial if the ordinary forms of treatment fail. Southey tubes and acupuncture are especially valuable in severe right heart failure with obstinate dependent edema. Care must be taken to avoid a severe low sodium syndrome with hyperkalemia.

- 5 Cool environment. Patients are usually more comfortable in a cool room.
- 6 Fluids. If sodium restriction is observed faithfully, there is no need for fluid restriction.
- 7 Exercise. Patients who are in bed should be given passive or active leg exercise to prevent thrombosis (see page 217).
- 8 Sedation. Patients with congestive failure may also experience insomnia. Use sedation carefully to ensure sleep.

D Therapeutic Myxedema. Useful in refractory resistant left ventricular failure, refractory anginal pain, uncontrolled ventricular rate in atrial fibrillation, and in frequent recurrences of atrial fibrillation not controlled with quinidine. It is successful in about 40% of well-chosen cases but is a potential hazard for the patient and should not be undertaken lightly. Myxedema is unpleasant and the cure may be worse than the disease.

Any of the measures used to treat hypertensive idioms may be employed (see page 370).

Observation During Treatment of Cardiac Failure

A careful clinical record should be kept of all patients being treated for cardiac failure so that their course can be fully followed and therapeutic changes made as needed. This record should include the following observations on every visit:

- 1 St tues of origi al symptoms
- 2 New sympto ns
- 3 Mo ning weight o w light w th s m lothes
- 4 P esen e of the signs of co g tiv f lur (veno engorge m nt and pulsatio s p lmo ary ale ple l fl id en gorg m t of th liver p ese of ed ms)
- 5 Examination of th he rt a d blood vessels (c diac sounds g llop hythm fr tio r b cardi rhythm and ap l rat ca diac at e pe iph ala te lai pul at on and stat of th veins)
- 6 Blo d pr u a d pr e c of pulsus alt nans

ACUTE PULMONARY EDEMA (cod No 324)

Acute pulmona y ed ma is a g s em rgen y. Prope under standing of th p ecip t ting f ctor d underlying d eas is n ces as y f r th phys ian to utilize the a io s therapeuti pos bilities i the individual cas (e g in s mild att ck morphin and est in b d alone m y s ffice m an att k d e to trial fib illation with rapid v nt c l rate la toa d C dig xin given intr enously may tak p ec d ce e some of the th r p o edur m tio d b low)

Eme g ncy T eatment

- A Pos t Th p tient should b elev ted to th semi Fowle b d po it on (see p ge 3) or plac d in a chair this de e s s th veno return to th h rt
- B M rphin Sulf t U S P B P 15 30 mg (1/4 1/2 gr) I V or I M Th anxiety d p es es es pulm n ry cfl es d i du es sleep Th att ndant l see ing of the for ful expir tion d eases the n g t ve i t alth a ic p ess e and the ven r turn to the h t
- C Oxygen wh n labl should be admini ter d i high con en f at ns Th is best ch ev d by mask or in the s of hild n by h od or t t Mode te on ent tion (40 to 60%) ca be s hie d by o ygen te t or asal thet Oxyg n re liev hypoxia d dysp ea and dec eas pulm nary capillary p rm ability (see page 144)
- D R d tio f Bl od Vol m
 - 1 Tou lo ta Soft rubber to n q ets o blood pre c lls pplied with ffici t p ess e to ob t ve us but n t a te lai flow d rot ted every 15 minutes will effec ti ly ed e the ou r t rn to the h rt Th tourniqu t sh uld b mo d g d ally a th atta k held s Appr x imately 700 f bl od m y be tr pped in the xtremities by this meth d
 - 2 V nes t on (300 to 700 cc) Th i the most dir t w y of r d i g the v t rn to th heart nd may tr ki gly l e the di o t p i and de ea th right at i l and p ripheral us p es r n l w tput di c f ilur Th p oedu should ot b do e if s emi is p t
- E R pid digitalization i f gr at val (see p g 197) E t m c e hould b cl d in g i ng digitali intra o ly to a p io sly digitali d p ti t

- F Aminophylline U S P B P 0 25 0 5 Gm (4 7½ gr) slowly I-V has been advocated. Oral aminophylline is relatively ineffective. 1 M aminophylline is often painful. Rectal aminophylline suppositories 0 25 0 5 Gm (4 7½ gr) may be helpful.
- G Hexamethonium In the acute pulmonary edema of hypertensive heart disease and in the presence of severe hypertension a slow intravenous infusion of hexamethonium 1 mg (¼60 gr) per minute to a dose of about 5 to 10 mg (¼12 ½ gr) may be very helpful. The infusion should be stopped when the systolic pressure falls to 170 mm Hg so as not to overstep the mark and produce hypotension.

REFRACTORY CARDIAC FAILURE

The refractory state is said to be present when the patient fails to obtain clinical improvement after the usual therapeutic measures outlined above. When this occurs the following procedure is advised:

- 1 Re-evaluate the total situation. Has the bed rest been adequate? Is the patient ingesting more sodium than ordered? Has he been receiving his therapy carefully? A review of the patient's activities, diet, and medications is essential.
- 2 Are unrecognized recurrent pulmonary infection, anemia, masked hyperthyroidism, vitamin deficiency, silent myocardial infarction, or arrhythmia present?
- 3 Have complications such as acute rheumatic myocarditis or subacute bacterial endocarditis been superimposed upon a rheumatic heart?
- 4 Are there electrolyte abnormalities which may have resulted from diuretics and resins if these have been used? Electrolyte disturbances may lead to mercurial resistance, produce a low sodium syndrome, or in the case of potassium enhance digitalis intoxication.

MANAGEMENT OF CONVALESCENCE

Specific Measures

- A Digitalis Once digitalis is instituted it is usually necessary to continue the administration of the drug for life (see page 185).
- B Low sodium Diet 1 5 Gm (22½ gr) NaCl (750 mg or 12 gr Na) per day (see page 53). It is advisable to check the patient's serum sodium or urinary sodium frequently to be certain that no deficiency is occurring. An inadequate sodium intake in the presence of severe renal impairment can precipitate fatal renal failure.
- C Mercurials These drugs should be used as frequently as indicated even as frequently as 2 to 3 times per week in prolonged maintenance in order to remove any accumulating edema (see page 204). Care must be taken not to cause sodium depletion or dehydration. Some patients will prefer the addition of salt to the diet as long as mercurials will remove it from the body. Many patients can be saved from severe chronic cardiac

invalidated by the liberal but judicious use of mercurials. Ammonium chloride may be given orally to potentiate the mercurials (see page 204)

General Measures

Provide adequate rest and exercise within tolerance. Careful attention should be paid to the treatment of the renal cardiac causes of cardiac failure (see page 182) and to the avoidance of precipitating factors (see page 182)

ELECTROLYTE DISTURBANCES IN CARDIAC FAILURE

During treatment of cardiac failure the type of electrolyte disturbance may be seen

Hypochloremia Alkalosis

A Mechanism Thus secondary to chloride excretion out of proportion to sodium loss following myocardial diuresis. This produces a low serum chloride and a high serum bicarbonate. Serum sodium and potassium levels may be normal or low. Symptoms of dehydration may be present: dry mucous membranes and loss of turgor and a latent or manifest tetany.

Treatment

1. Ammonium Chloride U.S.P. B.P. 4.6 Gm (1 1/2 dr) per day for 3 to 4 days and repeat after a rest interval of 3 to 4 days.
2. Potassium salts may be added if a deficit exists (see below). If tetany persists, calcium salt must be given concurrently (see page 302).

Low Sodium Syndrome

A Diagnosis The onset of weakness, oliguria, diaphoresis and at times headache, the low albuminemia. Heart with or without fever and vomiting are additional predisposing factors. Low serum sodium may be present without alkalosis or acidosis or it may be complicated by dehydration and acidosis. It may follow severe sodium restriction accompanied by mercurial diuresis.

Treatment

1. Mild cases. Increase sodium intake.
2. Severe cases. Treat with I.V. hypertonic saline (see page 15).

Hypokalemia

A This may result from excessive potassium excretion from mercurial diuresis or a total diuretic (Diurex®) or the use of a diuretic in a patient receiving a low sodium diet. Hypokalemia may induce digitalis toxicity and is manifested by muscular weakness, particularly of the muscles of respiration.

B Treatment Potassium Chloride U.S.P. B.P. 3.6 Gm (45.9 g) daily by mouth provided there is no renal failure. CAUTION: Potassium salts should not be given in the presence of a degree of renal failure.

PERICARDITIS

ACUTE FIBRINOUS PERICARDITIS (code No 420)

Acute fibrinous pericarditis may be caused by or associated with many diseases. The most common are rheumatic fever, uremia, collagen diseases, tuberculosis, viruses, and malignant disease. Acute fibrinous pericarditis generally produces little functional impairment for there is no mechanical interference with cardiac function. The most distressing symptom is pain and this may be entirely absent. It varies from local discomfort to very intense pain which is usually substernal or precordial and may be confused with angina or infarction.

Treatment

Treat the underlying condition and provide analgesics as necessary for relief of pain. Salicylates and/or corticotropin (ACTH) or the cortisones are useful in the nontoxic pericarditis (see page 518).

PERICARDITIS WITH EFFUSION (code No 420 100 8)

The diagnosis of pericarditis with effusion is important because the fluid accumulation may cause decreased cardiac output and venous return resulting in cardiac tamponade. This does not respond well to the usual treatment for cardiac failure (digitalis, low salt diets, etc.) but removal of the pericardial fluid may be lifesaving. This is infrequently required in the common varieties of pericarditis with effusion with the exception of tuberculous pericarditis. Both the rapidity of accumulation and the amount of fluid are important in determining the functional impairment.

Diagnosis

Symptoms and signs include apprehension, dyspnea, cyanosis, distended neck vein, tachycardia, pulsus paradoxus, increase of area of cardiac dullness, feeble or absent apex beat and diminished heart sounds. Pericardial friction rub may be present. The chest x-ray reveals a water bottle shaped heart shadow. The ECG shows low voltage of the QRS complex and T wave abnormalities. Diagnostic aids include cardiac catheterization or venous angiocardiography. Confirmation is by pericardial tap.

Treatment

A. Emergency Treatment (Paracentesis) The indications for pericardial paracentesis are the symptoms and signs of cardiac tamponade. As the pericardial fluid increases in amount and particularly when it increases rapidly the venous pressure may rise considerably and the cardiac output may progressively fall. When this occurs the patient becomes weak, pale, and dyspneic; the pulse pressure becomes very narrow and the pulse rapid and thready. If the patient goes into shock, under these circumstances the removal of the pericardial fluid may be lifesaving. The fluid should be removed slowly to avoid cardiac dilatation or serious cardiac reflexes.

1 Sites of puncture Avoid puncture of the ventricular muscle

- a Left 5th or 6th inter space about 1 cm within the area of cardiac dullness or 1.2 cm inside the left heart border sloped by x-ray (usually 7.8 cm outside of left paraxial line). The needle is pushed slowly inward and slightly upward. If effusion is present one should find fluid within 3.5 cm (at times 7.8 cm).
- b Epigastric area Area between xiphoid process and left costal margin. Insert the needle upward at an angle of about 30° and pointed toward the midline. The pericardium is reached about 3.4 cm.
- c Posterior approach To be used, as a rule, only when the above approaches are unsuccessful rarely used if one suspects pericarditis. On the 7th or 8th inter space in the mid-apical line. The left arm is elevated to rotate the scapula out of the way. The needle is directed inward and medially.

2 Equipment

- a No. 16 or 18 needle with short bevel and fitting stylet
- b No. 25 or 27 needle to infiltrate the skin with procaine
- c 20-30 cc syringe to remove fluid. Syringe should be connected to needle by a 4 inch piece of rubber tubing to prevent accidental movement of the needle.

3 Technique

- a Clean and sterilize skin over area to be punctured
 - b Draped surrounding area with sterile towels
 - c Infiltrate skin with 1-2% procaine solution
 - d Insert needle (detached from syringe and without a stylet) slowly into skin following directions according to site selected (above).
- Withdrawal of fluid When the fluid is aspirated it must be withdrawn very slowly. Sudden withdrawal of the fluid may result in acute cardiac dilatation leading to death. Some consider it advisable to replace half the amount of fluid withdrawn with air both to prevent excessive dilatation and to give better visualization of the process by x-ray. With the needle in place move 20 cc portions after the withdrawal of each portion injecting 10 cc of air.
- f After the needle is removed a simple bandage over the exit puncture is adequate.

B Specific Measures

- 1 Treatment of the patient (Code No. 420.123) The current treatment is to treat the systemic infection with bed rest, attention to nutrition, and other general factors as indicated by a fit bacteriologist. (p. 131) If the fever and signs of pericardial effusion do not rapidly subside and are still obvious in one month a gradual corticosteroid therapy should be considered in order to prevent chronic constrictive pericarditis. Judgment is required to determine when this is a step regardless of medical treatment and when signs of pericardial effusion appear.
- 2 Rheumatoid pericarditis with effusion (Code No. 420.198.8) Treatment for rheumatism. The salicylates may help in

190 Chronic Constrictive Pericarditis

causing fluid reabsorption. Paracentesis is usually unnecessary but should be performed if tamponade occurs.

- 3 Hydropericardium due to heart failure (code No 420 522 8) Treatment of the congestive failure is usually sufficient.
- 4 Hemopericardium due to rupture of adjacent structure (code No 420 532) Usually post traumatic. If fluid accumulation is excessive, remove fluid at once.

PURULENT PERICARDITIS (code No 420 100 2)

This is usually secondary to other infection elsewhere but is at times caused by contamination of a previous pericardial tap.

Treatment

A Specific Measures

- 1 Systemic chemotherapeutic agents. Treat infection with indicated chemotherapeutic agents (see page 514).
- 2 Interpericardial antibiotics. At the time of removal of the fluid, instill 50 000-150 000 units of penicillin or the approximate topical amount of streptomycin or other indicated antibiotic into the pericardial sac depending on organisms found (see page 514) and repeat whenever a tap is performed. Chemotherapeutic agents should be continued as long as purulent effusion is present.

B General Measures

- 1 Paracentesis. Perform as needed to relieve pressure.
- 2 Pericardiotomy. If fluid is encapsulated or patient is not responding to therapy, surgical drainage may be necessary.

CHRONIC CONSTRICTIVE PERICARDITIS (code No 420 100 4)

(Tuberculous pericarditis, code No 420 123 4)

This is usually due to tuberculous pericarditis. In the remainder of cases the etiology is unknown, a few instances occur following acute nonspecific pericarditis or traumatic pericarditis.

Treatment

A General Measures To combat ascites and congestive failure

- 1 Low sodium diet.
- 2 Mercurial diuretics as needed to keep patient dry (see page 204). This may be combined with intermittent ammonium chloride as in cardiac failure.
- 3 Digitalis is usually of little value.

B Surgical Removal of Constricting Pericardium This procedure can frequently restore a patient to normal health. If long-standing phenomena are chronic or the pericarditis is progressive, surgical intervention is the only method offering possible cure.

NEUROCIRCULATORY ASTHENIA (code No 004 580) (Da Costa's Syndrome or Effort Syndrome)

Neurocirculatory asthenia is a chronic disorder of young adults which is considered to be a psychiatric disorder. It is characterized by functional symptoms: dyspnea on effort, palpitations, left chest pain, and easy fatigability. The symptoms are often more closely related to the emotional connotations of effort than to the effort itself. Examination reveals no clinical findings of heart disease, although tachycardia is often present.

Treatment

- A Psychotherapy and Reassurance** The medical examination and the manner of handling the patient have important therapeutic value.
- 1 Medical examination should be thorough.
 - 2 The patient should be assured that no organic disease exists.
 - 3 Psychotherapy. Further and more intensive psychotherapy may be of value.
- B General Measures**
- 1 The treatment of hyperventilation. An acute attack may be aborted by the administration of 5% carbon dioxide rebreathing in a bag or by holding the breath. Do not give ammonium chloride. It does not relieve symptoms and may precipitate a crisis. Inasmuch as a fixed base has been lost, 3 mEq/kg of sodium bicarbonate is given.
 - 2 Good hygiene with moderation in all activities, a well-balanced diet, and proper sleep. Exercise is undertaken progressively when convenient.

Prognosis

The prognosis for survival is good but is often discouragingly poor for relief of symptoms.

PULMONARY HEART DISEASE (Cor Pulmonale)

Heart disease secondary to disease of the lungs of the pulmonary arteries. Emphysema, pulmonary fibrosis, silicosis, and kyphoscoliosis are common causes of chronic pulmonary disease.

Diagnosis

- A Symptoms** Symptoms of the underlying pulmonary disease may be present along with symptoms of pulmonary hypertension, preceded by many of the signs of pulmonary hypertension: right ventricular hypertrophy and failure.
- B Signs**
- 1 Signs of pulmonary hypertension. Systolic pulsation and murmur in the pulmonary area, a clear split second sound in pulmonary area, distant, clearly split second sound possibly with diastolic murmur of pulmonary insufficiency.
 - 2 Signs of right ventricular hypertrophy. Heaving right ventricular impulse in the left parasternal area, weak tapping precordial impulse, pulmonary gallop in the xiphoid area.

- 3 Signs of right ventricular failure follow usually with sinus rhythm and with arterial oxygen saturation less than 85%
- 4 Evidence of central cyanosis and high output state may be present

Treatment

A Specific Measures Appropriate antibiotic therapy for the respiratory infection that so commonly precedes failure in this type of case. The patient may be afebrile

B General Measures

- 1 Intermittent oxygen therapy possibly by positive pressure to increase arterial oxygen saturation and to decrease pulmonary arterial pressure. Continuous oxygen therapy should be avoided and the patient closely observed for stupor and coma since carbon dioxide retention may occur with oxygen therapy in these patients (see page 144). Intermittent positive pressure mask breathing e.g. with the Bennett or Emerson respirators at pressure settings of +10 to +15 (in inspiration) and 0 (expiration) may be helpful. The valve may be operated by an attendant if the patient does not breathe spontaneously. These devices provide a convenient effective method of administering bronchodilators, antifoaming agents and aerosols. They do not restrict cardiac output.
- 2 Incorporate intermittent positive pressure breathing is the single most effective therapeutic measure.
- 3 CNS depressants especially narcotics, barbiturates and hypnotics are strongly contraindicated in the treatment of cardiac failure secondary to primary pulmonary disease (cor pulmonale) due to their marked depressant action on the respiratory centers.
- 4 The usual methods of treatment of heart failure should be used (see page 182) bed rest, restriction of sodium, mercurial diuretics and digitalis. Digitalis may not be effective if there is a high cardiac output state.

CARDIOVASCULAR SYPHILIS

Cardiovascular lesions may manifest themselves as a complicated syphilitic aortitis, syphilitic aortic insufficiency, aortic or fusiform aneurysm of the aorta or as angiodysplasias due to involvement of the ostia of the coronary arteries.

The diagnosis may be supported by a history of syphilis, evidence of the disease elsewhere in the body (especially CNS syphilis) and a positive serological test for syphilis. Serological tests are negative in about 20% of cases.

Treatment

A Specific Measures

- 1 Treat latent syphilis (see page 440)
- 2 Therapeutic acidosis has the Herxheimer reaction rarely with penicillin. It is therefore not recommended. It is easy to precede penicillin treatment with iodides or bismuth.
- 3 Several subsequent courses of penicillin are advised by some authorities at intervals of 6 months or 1 year especially if the serology remains positive.

- B Postpone surgery for at least 3 weeks after recovery from congestive failure
- C Exercise caution when giving fluids containing sodium (including blood) to avoid producing pulmonary edema
- D Treat anemia prior to surgery
- E Treat malnutrition and avitaminosis especially avitaminosis B prior to surgery

THE CARDIAC PATIENT AND PREGNANCY

Status of Patient

The following information will assist in making an estimation of likelihood of cardiac failure

- A Function 1 class prior to pregnancy
- B Age of patient
- C Size of heart
- D Structural lesion of heart
- E Presence of arrhythmias
- F Socio economic status (e.g. if children are at home or if the patient must work)
- G Intelligence and cooperation of patient (e.g. can the patient rest? Can she stay on a low sodium diet?)
- H Presence of associated disease

Some Factors Which May Precipitate Failure in Heart Disease

- A Excessive work
- B Upper respiratory infection sodium retention
- C Anemia
- D Paroxysmal arrhythmia
- E Excessive sodium intake e.g. diet sodium bicarbonate for gastric acidity infusion of saline plasma or blood
- F Rheumatic activity

Assessment of Risk of Heart Disease in Pregnancy

- A Little or No Functional Impairment Practically all patients who are asymptomatic or who have only mild symptoms with ordinary activities may be allowed to continue to term under close medical supervision. If they develop more severe symptoms with activity they should be hospitalized and treated for failure and kept in bed until term.
- B Moderate or Marked Functional Impairment If the patient has pure mitral stenosis and develops acute pulmonary edema or has moderate to marked symptoms with activity mitral valvulotomy should be considered. This has been successfully accomplished up to the eighth month. If the patient does not have an operable lesion she should be hospitalized, treated for cardiac failure, and kept in bed until term.
- C Very Marked Functional Impairment All patients seen during the first trimester who have symptoms on little or no activity and who do not have an operable cardiac lesion should be aborted because of the high incidence of recurrent failure and death in this group of patients.
- D Sodium should be restricted after the second month.

Physiological Load When Pregnancy Imposes on the Heart

The work of the heart increases by about 50% at the beginning of both the third month when the blood volume and cardiac output increase. The placenta acts as an arteriovenous fistula. Cardiac failure may occur at any time from the end of the first trimester up to 2 to 3 weeks before term at which time the load from the placenta countable increases.

Management of Labor

- A Cesarean section holds that vaginal delivery is to be preferred except in those cases in which there is an obstetrical indication for cesarean section. Coarctation of the aorta may be the only exception to vaginal delivery because of the danger of rupture of the aorta.
- B The second stage should be made as short as possible using forceps when possible.
- C Ergonovine Maleate U.S.P. (Ergotrate®) should probably be avoided because of the increased work of the heart which follows.

CARDIOVASCULAR DRUGS

DIGITALIS AND DIGITALIS LIKE PREPARATIONS

Action of Digitalis and Digitalis like Preparations

- A It increases failure digitalis causes the reflex output of the myocardium thus increasing the efficiency of the heart. Digitalis significantly increases the diastolic pressure and in cardiac secretion of sodium and water and so establishes some of the hemodynamic abnormalities in the condition of cardiac failure. The increased diastolic pressure decreases the venous pressure however a direct effect on the venous motor system has also been postulated.
- B In the arrhythmias (especially atrial fibrillation and flutter) digitalis slows conduction between the atrium and ventricle as demonstrated by the S-A and A-V node both by direct action and by stimulation of the vagus nerve.

Principles of Administration

- A Concept of Digitalis Saturation (Digitalization) Digitalis must be administered initially in a large dose in order to achieve tissue saturation and obtain a therapeutic effect. After digitalization has been accomplished the smaller dose prescribing the amount metabolized and excreted is administered daily a long indication of digitalis per day (usually for life).
- B Criteria of Adequate Digitalization Digitalis is administered until a therapeutic effect has been obtained (e.g. relief of congestive failure or slowing of the ventricular rate in atrial fibrillation) or the earliest toxic effect (anorexia) is observed. Congestive failure with normal rhythm. Diuresis is indicated if edema fluid is present.

- b Cardiac size is decreased as dilatation becomes less
- c Venous pressure and circulation time return to normal
- d Decrease of heart rate results if increase was due to failure
- e Engorged tender liver becomes smaller and non tender

2 Atrial fibrillation When the rate is below 80 after exercise one can usually consider the patient adequately digitalized. The following simple exercises are adequate

- a Bed patients Sit up 5 times
- b Ambulatory patients - Hop up and down on one foot 5 times

3 Ecg effects The most characteristic change which digitalis produces in the Ecg is sagging of the ST segment and displacement of the T waves in a direction opposite to that of the main deflection. Later there may be a prolonged P R interval. The ST T changes cannot be used as criteria of digitalis toxicity for the effects appear before saturation is present and persist for 2 to 3 weeks after digitalis has been discontinued. However the Ecg is often of value in determining whether digitalis had been administered in the past 2 to 3 weeks and may give one an idea of the amount.

C Toxic Effects of Digitalis There are no non toxic digitalis preparations and the difference between the therapeutic and toxic level is very small

- 1 Slight toxicity Anorexia
- 2 Moderate toxicity Nausea and vomiting headache malaise
- 3 Considerable toxicity Diarrhea ectopic beats (especially ventricular) blurring of vision confusion disorientation
- 4 Gross toxicity Severe diarrhea abdominal pain high degree conduction blocks and atrial or ventricular fibrillation

D Relationship of Digitalis to Potassium Ion Recent work indicates that there is an antagonism between potassium and digitalis and that digitalis toxicity is more likely to occur in any clinical disorder in which decreased potassium of the cell or of the serum is present. This may occur with potassium deficiency from mercurial compound following cortisone therapy or in any condition in which deficiency of serum potassium is produced. In these circumstances the dose of digitalis should be reduced or potassium should be given.

E Treatment of Severe Digitalis Intoxication Omit digitalis completely until the intoxication has subsided and treat the cardiac failure with other means. Give potassium salt 4 to 8 Gm (60 to 120 gr) orally per day in divided doses or depending upon the clinical urgency well diluted intravenous potassium salts slowly (not more than 10 to 20 mEq/hour).

The differentiation of digitalis toxicity and delayed digitalization is sometimes quite difficult. The only safe procedure if uncertain is to omit digitalis and treat the cardiac failure with resuscitation of sodium mercurial diuretics and other means to improve the cardiac function. If one uses vomiting or arrhythmias which are suspected of being due to digitalis are in fact due to digitalis they will subside in 2 to 3 days if digitalis is stopped. The use of rapid acting intravenous

digital preparations tend to differentiate these two clinical states is not advised because of the risk involved

- F The cardinal principles of digitalis therapy remain the same whether one uses a crude drug such as the whole leaf or one of the purified glycosides. They all have broadly similar pharmacological actions differing only in the extent of the therapeutic effect. The differences may be utilized to advantage in the treatment of the individual patient particularly with respect to potency speed of action extent of absorption and duration of action.

Indications for Administration

A Indications for Administration of Digitalis

- 1 Cardiac failure or combined with the other sinus rhythm or atrial fibrillation
- 2 Atrial fibrillation or flutter with a rapid ventricular rate
- 3 Supraventricular paroxysmal tachycardia
- 4 Pre-operative especially mitral valvulotomy in patients with sinus rhythm so that if paroxysmal atrial fibrillation occurs during or following surgery the ventricular rate will not be excessively rapid
- 5 The prevention of paroxysmal atrial rhythms in patients in whom quinidine has failed or cannot be tolerated

B Risks of Administration

- 1 Overall administration is seldom necessary when verapamil is needed and prevents administration of digitalis
- 2 Potential administration
 - Emergence of digitalis toxicity
 - (1) Atelectasis, pulmonary oedema, other effects

Caution should be used in giving the full digitalizing dose in a single injection especially under these circumstances. The drug should be given slowly in divided doses.
 - (2) Treatment of atrial rhythms when then needed for control of the ventricular rate is urgent
- b Inability to take digitalis orally. Nausea and vomiting due to any cause, or a postoperative patient.

Method of Digitalization

A Urgent Cases When the patient has lived a digitalis preparation in the preceding 2 weeks

- 1 Parenteral digitalization (indicated) **CAUTION**
Never administer a full digitalizing dose I.V. unless it is certain that no digitalis has been taken in the preceding two weeks. Always give I.V. preparations slowly.
 - a Select drug depending on the rapidity of effect needed
 - b Initial dosage schedule. Except in major emergency do not give the average digitalizing dose in a single dose. A good general rule is to give $\frac{1}{2}$ to $\frac{2}{3}$ of the average digitalizing dose immediately and follow with the remainder in 2 to 4 hours. Observe fully for digitalis toxicity (page 198).

Addition of oral digitalis. At the time that the initial dose is given parenterally it is advisable to give orally

DIGITALIS AND DIGITALIS LIKE PREPARATIONS

Qty id	P p tion A il bl	Digitalis D	M th d r Administ ti	Sp d f Ma Intern A tion and Dur tion	Maint nanc D
Ovalb U S P B P	1 amp 1 0.25 mg (1/240 gr)	0.25 0.5 mg (1/240 1/20 gr)	0.25 0.5 mg (1.2) dil 1 d in 10 all el wly I V flow with anoth d g (b low)	1/2 1 1/2 h d tion 2 4 d y	Not u ed f maint anc
De la oaid U S P (C di nld D ⁸)	2 and 4 amp 1 0.4 and 0.8 mg (1/250 d 1/75 gr)	8 (1.6 mg) (1/40 gr)	6 (1.2 mg) I V I M and f ll w by 1 2 (0.2 0.4 mg) I V I M q 3 4 hour till f l s obt i d	1 1 1/2 d r tion 3 6 d y	1 2 (0.2 0.4 mg) (1/200 1/50 gr)
U S P (Dilut b f)	1 and 2 amp 1 0.2 0.4 mg (1/200 and 1/50 gr)	1.2 mg (6) (1/50 gr)	0.6 mg (3) I V I M f ll w d by 0.2 0.4 mg q 4 6 hours till 1 2 mg i g l	3 8 1 d ti n 14 21 d y	0.05 0.2 mg (1/200 1/50 gr)
Dig in U S P B P (Dilut b f)	1 amp 1 0.5 mg (1/20 gr)	1.5 mg (3) (1/40 gr)	1 mg (2) (1/20 gr) I V 2 d 0.5 mg (1) (1/20 gr) in 3 4 h 2 th n 0.25 mg (0.5) (1/240 gr) q 3 4 hours until fte t to btain d	1 2 h d att 3 6 d y	0.25 0.75 mg (0.5 1.5 c) (1/240 1/80 gr)
Digitalis U S P B P	0.03 0.05 and 0.1 Gm tabl t (1/2 1 and 1 1/2 g)	1 0 1.5 Gm (13 22 1/2 gr)	0.8 Gm (10 g) t 0.4 Gm (5 g) in 8 h 0.2 Gm (3 g) q 6 h r 2 3 d th n 0.1 Gm (1 1/2 g) b 1 d til fct i br i d	6 8 h d tion 18 21 day	0.05 0.2 Gm (3/4 3 0 gr)
Digitalis U S P	0.1 0.2 mg t bl t (1/20 1/50 gr)	1.2 mg (1/50 gr)	0.6 mg (1/20 gr) t and ep t in 12 h on and th 0.2 mg (1/20 gr) b 1 d til fct i br i d	6 8 h d ti n 14 21 d y	0.05 0.2 mg (1/200 1/50 gr)
Dig in U S P B P	0.25 0.5 and 1.0 g tbl t (1/240 1/20 d 180 gr)	2.5 mg (1/20 1/20 gr)	1 mg (1/20 gr) t d th q 0.5 0.75 mg (1/20 1/20 gr) q 6 h T 1.8 mg (1/20 gr)	4 8 1 d ti 2 6 d y	0.15 0.50 mg (1/400 1/20 gr)
Lan 1 id C N P (C di nld ⁸)	0.5 mg tabl t (1/20 g)	7.5 mg	3.5 mg (1/20 gr) t 1 mg i 8 th 0.5 mg q 6 h until fct i br i d		0.5 2.5 mg (1/20 1/25 g)

PARENTERAL

a erage maintenance dose of the preparation used. If the patient is able to swallow. By instituting the drug orally, optimum digitalization can be achieved and maintained from the start. It is not necessary to give the same digitalis glycoside orally as that used for the initial medication (e.g., many digitalis with I.V. lanatoside C and give digitalis folium for maintenance).

- d. Caution must be exercised. A previous history of digitalis therapy is often difficult to obtain because the new preparation is tasteless and may be in tablets of various colors. The patient may be unaware of the therapy he has been receiving.

Digital toxicity has been seen in patients who have denied or were unaware of having received the drug. This is another reason for avoiding a full digitalizing dose in a single injection.

- e. Individualize course of each patient. No dosage schedule will fit all patients.

2. Rapid oral digitalization (within 24 hours). A single oral digitalizing dose is usually unwisely since nausea and vomiting are common making stimulation of degree of digitalization very difficult.

Multiple oral doses are usually quite adequate and rapid if carefully followed. In all cases, close medical supervision is required for each dose. As digitalization is approached, the drug should be stopped at the first sign or symptom of toxicity (p. 198).

Oral Administration of the Digitalis Drugs

Usage	Drug	How Administered
Maintenance	Digitalis	0.4 Gm (6 gr) q 8 hours for 3 doses
	Digitalin	0.4 mg (1/150 gr) q 8 hours for 3 doses
	Digitalin	1.0 mg (1/60 gr) q 8 hours for 3 doses
Intermediate	Digitalis	0.2 Gm (3 gr) tid for 2 days
	Digitalin	0.1 Gm (1 1/2 gr) qid for 3 days
	Digitalin	0.2 mg (1/500 gr) tid for 2 days
Late	Digitalis	0.5-0.75 mg (1/120-1/80 gr) tid for 2 days
	Digitalin	0.25-0.5 mg (1/240-1/120 gr) qid for 3 days
	Digitalin	0.1 Gm (1 1/2 gr) tid for 4-5 days
Late	Digitalin	0.1 mg (1/600 gr) tid for 4-6 days
	Digitalin	0.25-0.5 mg (1/240-1/120 gr) tid for 4-6 days

3. Slow digitalization. At times it is difficult to digitalize slowly over the few weeks, especially if the patient cannot tolerate the oral dose. If this is the case, any of the preparations can be given in daily doses of 2 to 3 Gm with a tapering maintenance dose of 1 to 3 Gm. The total digitalizing dose may be somewhat greater than when rapid digitalization is accomplished. One must individualize

schedule is maintained. If high blood levels are desired the individual dose must be increased so the interval between doses is shortened.

Because of the fact that 30-40% of the peak blood level of quinidin is still present in the serum 12 hours following a repeated dose of quinidine a fixed dosage schedule such as 0.4 Gm (6 gr) every 2 hours for 5 doses can be repeated for several days to produce a certain concentration of quinidin in the blood.

U

Widely different opinions have been expressed by various cardiologists on the indications, dosages and dangers of the use of quinidin. It must be remembered that patients in whom quinidin has been used have organic diseases and a pediatric consultant occur even when quinidin is not given to these individuals. Until recently most of the symptoms of blood quinidin detoxification has been available the symptoms of the effect was often not an arbitrary rather than a quantitative basis.

A Indications

- 1 Ventricular tachycardia
- 2 Conversion of atrial fibrillation to a normal rhythm. Most diologists feel that the presence of marked cardiac failure is a contraindication as digitalis is indicated for the treatment of quinidine.
- 3 Atrial fibrillation if digitalis fails to produce sinus rhythm.
- 4 Paroxysmal atrial and nodal tachycardia.
- 5 Prevention of re-entrant paroxysmal rhythms.
- 6 Suppression of frequent premature beats peculiarly following myocardial infarction. Stoppage of the treatment.

B Contraindications

- 1 Idiopathic syncope manifested by frequent paroxysmal attacks are hypotension following the treatment of 0.1 Gm.
- 2 Complete heart block } R.I.T.
- 3 Bundle branch block } contraindications
- 4 Thyrotoxicosis
- 5 Acute rheumatic fever
- 6 Subacute bacterial endocarditis

Relative Adverse Effects

A Oral (Quinidine Sulfate, U.S.P., B.P.) This is the most effective route with parenteral quinidine as particularly indicated (See page 203 for dosage).

B Parenteral

- 1 Intramuscular preparations. The intramuscular preparation can be used if the patient is unable to take the medication orally and the situation not critical.
Quinidin Gluconate N.F. 0.8 Gm (12 gr) in 10 cc ampules.
- 2 20% Quinidine Sulfate U.S.P. B.P. in propyl glycol.
15% Quinidin hydrochloride dissolved in water and tyrosine.
- 2 Intravenous preparation. An intravenous preparation should be used only when great urgency requires and only by

and no dose will fit all patients. The patient should be instructed regarding the early toxic symptoms when they occur the drug should be stopped for one day and the patient then given the average maintenance dose.

- B Partially Treated Cases If a digitalis preparation has been taken within 2 weeks give $\frac{1}{4}$ of the estimated digitalizing dose and then give additional digitalis cautiously observing patient's response.

Maintenance Dosages and Methods

The oral route is preferred in maintaining digitalization. The exact maintenance dose must be determined clinically for each patient. (The table on page 199 gives the average doses.)

QUINIDINE

Quinidine is a valuable drug in the management of most cardiac arrhythmias. Quinine may be used but is only about 30% as effective as quinidine. Only quinidine will be discussed here.

Pharmacology

- A Action Knowledge of the pharmacological effects of quinidine is important in order to understand the use of the drug. Quinidine has a variety of actions:

- 1 It increases the refractory period of cardiac muscle.
- 2 It slows the rate of atrial and ventricular conduction.
- 3 It decreases the excitability of the myocardium.
- 4 It reduces vagal tone.
- 5 It is a general depressant to smooth muscle.

As far as conversion of atrial fibrillation is concerned several of these pharmacological actions oppose each other. The clinical effect depends on which of the effects predominate.

B Clinical Pharmacology

- 1 Route Can be given orally I.M. or I.V. as occasion demands. The I.V. route should be used only by those experienced in the use of the drug and no gentler titration.
- 2 Absorption Orally quinidine is rapidly absorbed, reaches a peak level in about 2 hours and is relatively slowly excreted. There is a slow elimination to about 30% of the peak level after 12 hours.
- 3 Excretion and fate Only 10-20% of orally administered quinidine is excreted in the urine; the remainder is metabolized in the body.
- 4 Dose per day After the same dose of the drug is continued for 5 or 6 days at 2-hour intervals no significant rise in blood level occurs with further doses at the same interval.
- 5 Cumulative effect When a fixed dose of quinidine is given 4 times a day as in a maintenance schedule the blood level rises progressively but more slowly, reaching a maximum in about 48 to 72 hours. The midday blood level then remains more or less the same as long as this method is

- C If the patient has atrial fibrillation or atrial flutter, complete digitalization is advised to lower the ventricular rate and to improve cardiac function. If digitalis is not used, the decreased A-V conduction ultimately for quinidine may cause a sinus bradycardia rate of 30-50 beats per minute and may force cessation of quinidine therapy.
- D If cardiac failure is present in a patient with chronic hyaline arteriosclerosis, sodium restriction and diuretics (etc.) should be used prior to quinidine therapy. A period of complete ambulatory status following the treatment of cardiac failure is also advisable to decrease the likelihood of venous thrombosis. Anticoagulant may be desirable for a week prior to quinidine.
- E A satisfactory method of quinidine administration is as follows:
 0.4 Gm (8 g) every 2 hours for 5-6 doses on the first day.
 This produces an average blood level of 6-7 mg/L. Each succeeding dose produces a progressively smaller increment in the blood level and, if necessary, do not start after 5-6 days a high dose of the drug must be instituted. If the situation is urgent this may be started after the fifth dose or one may wait until the next morning and start the dose with 0.6 Gm (9 g) every 2 hours. Giving the drug more frequently than every 2 hours does not permit the peak effect of the preceding dose to be reached. In most cases 0.6 Gm (9 g) every 2 hours for 5 doses will convert the arrhythmia to sinus rhythm. If not high doses can be used if no toxicity has been encountered and it is urgent to convert the rhythm. Eighty percent of the successful conversions occur with daily doses of 3 Gm (45 g) or less. If doses greater than this are used, successful conversion will be less likely and toxicity greatly increased. Quinidine effects on the myocardium by:
 1. Progresses to increase in blood quinidine level.
 2. Determining that of the fibrillatory wave is obtained by V_1 , the right precordial lead, the electrocardiogram. The atrial rate slowed markedly; atrial fibrillation is the indication of the 200-250/min conversion is seen.
 3. Measurement of the QT interval and QRS complex. As the measurement increases up to 25-30% above the initial values, significant differences are noted.

NITRITES

The nitrite acts by relaxing the smooth muscle, especially of the coronary arteries and also of the other smaller blood vessels as well as in blood pressure. The rapid-acting nitrites relax the angiotensin II, slowing the drug may benefit a few selected cases of hypertensive emergency occasionally by its action on the peripheral vasculature.

physician familiar with the use of the drug Quinidine Gluconate N F 0.8 Gm (12 gr) in 10 cc ampules can be diluted with 50-100 cc 5% glucose and given slowly I V at 1 cc per minute

Toxicity

A Idiosyncrasy See page 201

B Toxic Effect

- 1 The myocardial toxicity is the most important and should be specifically looked for when quinidine is used. The earliest effects are seen electrocardiographically
 - a Prolongation of the Q-T interval
 - b Prolongation of the QRS interval
 - c Ventricular premature beats or ventricular tachycardia
- 2 Nausea, vomiting and diarrhea. These are rarely critical but may be sufficiently severe to require cessation of the drug.
- 3 Cinchonism. Tinnitus, vertigo and headache are usually mild but may be important enough to require stopping the drug.

CAUTION When the QRS interval becomes more than 50% wider than that seen before treatment or when runs of ventricular premature beats or ventricular tachycardia occur quinidine should be stopped immediately. In patients with atrial fibrillation who are converted with quinidine transient S-A block may occur at the time of conversion and nodal rhythm may be temporarily noted. This has not proved to be of clinical significance. In very rare instances ventricular tachycardia may progress to ventricular fibrillation and sudden death. Prolongation of the P-R interval is occasionally seen for a short time when sinus rhythm follows quinidine conversion of atrial fibrillation. This is rarely serious and usually subsides spontaneously as the smaller maintenance doses of quinidine are employed.

- 4 Other cardiovascular effects
 - a Hypotension may occur when large doses of quinidine are used or if the drug is given parenterally. It rarely is significant with ordinary oral dose.
 - b Embolic phenomena. Emboli occur in about 1% of patients with chronic atrial fibrillation converted with quinidine. The incidence is higher in untreated atrial fibrillation. In fact atrial fibrillation with frequent emboli is an important reason to attempt conversion to sinus rhythm. Anti-coagulants are advised for 1-2 weeks prior to conversion in these cases to prevent the development of new thrombi in the atria. The hazard of emboli with quinidine has been exaggerated but must be appreciated and regarded as a calculated risk.

Procedure for Conversion of an Arrhythmia to a Normal Rhythm

- A The patient should be under constant observation preferably in the hospital where frequent examination of apical cardiac rates and electrocardiograms may be taken.
- B A test dose of 0.1 Gm (1 1/2 gr) has been sufficiently used to exclude possible idiosyncrasies. Wait 2 hours.

usually if the patient on a low sodium diet inc the
g t may cause od m depletion o hypo hlor mic alkalosi (s e
page 187)

Parental Preparations

- 1 M u phylline Injecti n U S P (M cuza thin[®]) 1.2 cc
I V s n e d d
- 2 Mersalyl and Theophyllin Inject on U S P Inj tion of
Mersalyl B P (Saly ga Theophyllin[®]) 1.2 c I V as
d d
- 3 M all id l j tion U S P (M uhydrin[®]) 1 I V
o l M s n e e d d
- 4 M e c p t o m e r n S o d u m U S P (Thi meri S o d u m[®]) p
p l i e d a s d r y p o w d e r i n v i a l 1.4 Gm (21 gr) i n 10 c
i a l 4.2 Gm (63 g) i n 30 a l A d d d i s t i l l e d w a t e r t o
b r i n g t o p e r v u l u m a d r f i g r t e G i v e 0.5 t o 2 c c
s b c t a s n e e d e d M y a l s o u s i M

Oral Preparation

Alth gh th o a l p e p a t i o n a r e n o t f u l l y e v a l u a t e d a n d m y
b a t i c i n t h e i t i t h u e m a y a t t i m b e w r a n t e d
S e e i r e l l y a v a i l a b l e

- 1 M a l l i d I n j t i o n U S P (M e r u h y d r i n[®]) w i t h A o b i
A c d U S P 1.2 t b l e t s a f t e r v e r y m a l
- 2 C h l r m o d n N N D (N o h y d n[®]) 18.3 mg (10 mg
H g) 1 t a b l e t o r m o r e d a i l y a s n e e d e d

OTHER DIURETICS

- 1 A m m o n i u m C h l o r i d e U S P B P 4.6 Gm (60.90 gr)
d i l y f o 3-4 d a y f o l l o w e d b y a r s t p e r i o d o f s m i l a d
t i o n U s f u l a o a s a p o t e n t i a t i n g a g t f o t h e a t i o n o f
m c i s
C b o n a n h y d r a h b t o A i s l m d e N N D
(D m o x[®]) 250-500 mg (4 7 1/2 g) d y 2 t o 3 d a y
w e k f p t u l a v l i h o p i t o r y a c d o s i s
3 C h l o t h i d (D i l e[®]) 250 mg (4 g) 3 t i m e s d a i l y
a w p o t t l d t T h a v a l a b l d t a s e s c t y
b i t h d u g p p e t o b e a p t t a t h m e r i
p o d u g o d m d s i s

PROCAINAMIDE HYDROCHLORIDE U S P (PRONESTYL[®])

P o a m i d d p s t o p i p m k s p v t s a h y t h
m i a u n d e c y l p p a s t h e i a f o l l o w i g e p i p h d i
f u l l i t h t i m t o f o d a l a n d v t i u l a a h y t h m i a T o a
l e s d g e t b e d t o p v t h a r h y t h m i a s I t h e
n h l p t t e f f t o t h e a t i l t h n n t h v t r i u l a r
h y t h m i C l i n i l x p i e c i s t i l l t o o l i m i t d t o s t a t e w h t h e
p o c i m d q u i d u n i a t h d r g f h o l e i n t h e v e t r i c u l a
h y t h m i s

Nitrite Preparations

Preparations and Dose	How Administered	Speed of Action and Duration
Amyl Nitrite U S P B P Pearl contains 0.2 cc (3 n)	Break pearl in cloth inhale p r n	Onset 10 sec Lasts 5-10 min
Glyceryl Trinitrate Tablets U S P B P (Nitroglycerin) 0.3-0.6 mg ($\frac{1}{200}$ - $\frac{1}{100}$ gr)	1 tablet placed under tongue p r n	Onset 1-2 min Lasts 15-40 min
Pentaerythritol Tetranitrate N N D (Peritrate [®]) 10 mg ($\frac{1}{6}$ gr) tablets	Orally every 4-6 hours before meals	Onset 15-30 min Lasts 4-6 hours
Sodium Nitrite U S P B P 30-60 mg ($\frac{1}{2}$ - 1 gr)	Orally every 3-4 hours	Onset 5-10 min Lasts 1-2 hours
Erythrityl Tetranitrate Tablets N F 15-30-60 mg ($\frac{1}{4}$ - $\frac{1}{2}$ - 1 gr)	Orally every 4-6 hours	Onset 15-20 min Lasts 3 hours
Mannitol Hexanitrate Tablets N N D 15-60 mg ($\frac{1}{4}$ - 1 gr)	Orally every 4-8 hours	Onset 15-30 min Lasts 4-6 hours

XANTHINES

Recent studies with cardiac catheterization and metabolic balance studies have demonstrated that xanthines increase the cardiac output, increase renal blood flow and glomerular filtration rate, and enhance the excretion of sodium and water. They therefore may be valuable in the treatment of cardiac failure. In addition, they have been shown to increase the coronary blood flow when used in large doses and may on occasion be helpful in angina pectoris.

Preparations

- A Oral A variety of official preparations are available but a satisfactory one is Aminophylline U S P B P (enteric coated) 0.1-0.2 Gm ($\frac{1}{2}$ - 3 gr) 4-6 times per day.
- B Parenteral Aminophylline Injection U S P B P 0.25-0.5 Gm ($\frac{3}{4}$ - $1\frac{1}{2}$ gr) 1 V slowly over a 5 minute period 0.1 M may repeat in 2-4 hours.
- C Rectal suppositories containing Aminophylline U S P B P 0.36-0.5 Gm ($\frac{5}{8}$ - $1\frac{1}{2}$ gr) may be valuable in an impending attack of cardiac asthma or in nocturnal angina pectoris.

MERCURIAL DIURETICS

The mercurial diuretics act by reducing the tubular reabsorption of sodium and chloride. They may be used for edema due to most causes except those associated with impaired renal function. They are of great importance in congestive failure. A o d e x e a i e

Chapter 8

DISEASES OF THE BLOOD VESSELS

PERIPHERAL ARTERIAL DISEASE

An important consideration in the management of patients with peripheral arterial disease is the determination of (1) the amount of disability due to spasm and (2) the amount of disability due to occlusion. Therapy is aimed in each case at relieving these disturbances.

Differential Diagnosis of Common Peripheral Vascular Diseases

	Rhyndaud's Disease (code N 47x 502)	Thromboangiitis Obliterans (code No 40 930)	Arteriosclerosis Obliterans (code No 460 952)
Sex	70-90% female	97% male	Over 75% male
Age at onset	10-50 years	20-35 years	Over 40 years
Extremities involved	Usually upper but may occur lower	40% upper extremities 98% in lower	Always lower rarely upper
Symmetry	Symmetrical bilateral	Asymmetrical usually bilateral	Asymmetrical usually bilateral
Peripheral arterial pulsation	Present	Absent or diminished	Absent or diminished
Usual sites of gangrene	Small toes tips of fingers and toes	Variable	Variable
Venous involvement (phlebitis)	Absent	Often present	Absent
Calcification in arteries	Absent	Rare	Usually present

Degrees of Spasm and Occlusion in Peripheral Vascular Diseases

Disease	Spasm	Occlusion
Arteriosclerosis obliterans	0 to +	+++
Thromboangiitis obliterans (Berg's disease)	++	++
Rhyndaud's disease	+++	0 to +
Autointoxication	+++	+++

Dosage and Administration

- A Oral Preparation (250 mg capsules) 0.25 to 1 Gm (4-15 gr) orally every 4-6 hours is the recommended dose.
- B Intramuscular Preparation (1 Gm ampules in 10 cc diluent) The peak effect occurs within 15-60 minutes and a significant blood level is still present after 6 hours. The blood level is higher and the decrease is slower in patients with congestive failure and renal insufficiency. Hypotension is infrequent with the intramuscular use of the drug in the above dosage.
- C Intravenous Preparation (1 Gm ampules in 10 cc diluent) Can be used for ventricular tachycardia of a severe or urgent nature. The drug should be given very slowly 50-100 mg ($\frac{3}{4}$ -1 $\frac{1}{2}$ gr) per minute up to a dose of 1 Gm (15 gr) with continuous blood pressure and if possible electrocardiographic control.

Toxicity

The same precautionary methods outlined in the sections dealing with quinidine are essential when procainamide is being used.

- A Severe Hypotension This is noted particularly with the parenteral use of procainamide and may be severe enough to require cessation of the drug. This is why frequent blood pressure demonstrations are necessary while the drug is being given.
- B Conduction Defects Prolongation of the QRS interval may occur as with quinidine.
- C Ventricular arrhythmias may occur as with quinidine.

A General

- 1 Correlate any disorder (e.g., angostiv faller) which may interfere with the blood supply
- 2 Disabilities present must be vigorously controlled
- 3 Tobacco in any form should probably be prohibited but there is no complete agreement on this point except in thromboangiitis obliterans or Buerger's disease where treatment is usually in the patient with continuous smoking
- 4 Alcohol beverages in moderation are not contraindicated
- 5 A well-balanced nutritious diet should be maintained
- 6 Adequate rest and relaxation avoid fatigue

B Local Measures

- 1 Avoid extremes of heat and cold do not use contrast baths
- 2 Fingernails to a free fingertip controlled Fungicidal dressings to (e.g., Castles) may be used Avoid using Whitfield's ointment (see page 89)
- 3 Infection of and trauma to the affected extremity must be guarded against. The patient should be given the following instructions:
 - a. Soak feet for 10-15 minutes in warm water (not hot water) before dressing
 - b. Bunions or corns must be treated by physician or a chiropodist
 - c. Skin must be kept soft and pliant by rubbing with lanolin or a bland vegetable oil 2 times daily
 - d. Socks should be changed at least once a day. For warmth use soft woolen socks. 2 pairs of another kind. Shoes must be well fitted and properly supported

C Special Measures The following may be used in an attempt to increase circulation

1. B. G. exercises may be of value. However, do not if an infection or open wound present. Individualize the exercises. Demonstrate and then refer to the physical therapist.
 - a. Elevate leg about 45 degrees (suppose the man is inverted) above the level of the heart. Hang or painless (usually 15 minutes or less)
 - b. Next allow the leg to drop freely for 2-5 minutes until maximum numbness. At the same time the feet are moved downward and upward and then downward and upward. The toes are spread apart and while the movement is being made. Do the opposite exercise 10 times. If the feet are painful it may be necessary to limit the exercise. The plantar flexion in horizontal position for 2 minutes
 - c. Repeat the complete exercise 5 times each session and have 3-5 sessions daily
2. Manual massage may be substituted. Probable that the only effect is to increase the circulation
3. Vasodilator drugs (page 211). The cardiovascular system of little or no value and unless there is a normal vasoconstriction may actually be harmful. Blood flow studies show a decrease in the blood supply to the ischemic limbs if the locally administered vasodilator the height of systemic vasodilation due to drug

Differentiation of Spasm and Occlusion

	Spasm	Occlusion
Color	Livid cyanosis	Blanched
Moisture	Wet	Dry
Veins	Constricted	Full dilated
Temperature	Cold	Cold
Reaction to vasodilating tests	Extremity becomes warm	Extremity remains cold

Adequate differentiation can usually be made on the basis of the first 3 of the above factors. Peripheral arterial disease usually is a mixture of spasm and occlusion but in many cases one factor is more prominent than the other. Therapy is aimed at correcting the physiological abnormalities whenever possible.

Test for Degree of Arterial Occlusion

A simple technique for evaluating the degree of arterial occlusion in the lower extremities especially the foot is the reactive hyperemia adelson test. The test is particularly useful in evaluating treatment and in determining the prognosis of the foot.

A. Technique

1. The patient is placed supine and the brachial blood pressure taken.
2. The toes are raised to 65 cm above the actual level and observed for blanching. (The actual level is taken at 7 cm below the junction of the manubrium and the body of the sternum (angle of Louis)).
3. If no blanching occurs the feet remain elevated and blood pressure cuffs are inflated just above the ankles to a pressure 50 cm above brachial systolic pressure. The occlusion cuffs are left on for 5 minutes.
4. At the end of that time with the feet still elevated the pressure in the cuffs is suddenly released and the feet observed for return of color.
5. If at the end of 1 minute color has not returned the foot is lowered 5 cm and then lowered 5 cm every 30 seconds until color returns. The level at which color returns is noted.

B. Interpretation

1. If the filling pressure (level at which color returns) is 35 cm or more above the aural spontaneous healing of an ulcer will occur or if amputation is necessary through the foot the amputation site will heal.
2. If the filling pressure is under 35 cm the more extensive procedures (e.g. sympathectomy, endarterectomy) or drug therapy must be done to help raise the pressure.

CHRONIC OCCLUSIVE ARTERIAL DISEASE (Usually Arteriosclerosis)

Treatment

Primarily conservative but thromboendarterectomy, vascular grafts and sympathectomy are of inestimable value in the properly selected case.

Criteria for sympathetic B st determined on clinical grounds assisted by vasodilator test

(1) Clinical evidence of increased vasomotor tone. This is evidenced by swelling (cyanosis and congested veins (absence of severe rubor and no malodorous blanching reaction on elevation)

(2) Sympathetic block on ml test gives relief of pain and intermittent claudication and better color to feet

b. Clinical findings

(1) Marked rubor on dependency

(2) Rapid blanching on elevation

(3) Atrophy of tissue

* Vasodilator drugs

a. Chemical sympathectomy. The introduction of ganglionic blocking agent has afforded a new approach to the relief of abnormal vasoconstriction. Many have been tried but only few are useful.

(1) Adrenergic blocking agents. These drugs act either by binding in the vascular musculature (e.g. of the foot) which is probably the most desirable mode of action. They thus not only block the sympathetic vasoconstrictor stimuli but also the vasoconstrictor effects from circulating epinephrine and norepinephrine. This group of drugs includes Tolazoline, Hydrochloride, LSP (Pascoline®), phoxybenzamine (Dibyl®), Azepate, Phosphate, NND (Iliad®) and Nylid in Hydrochloride, NND (Aridin®) (table). These drugs are effective orally. They are thus useful in the acute and subacute malvasospasm.

Side effects which are tolerable but of serious magnitude are gastrointestinal spasms, spasm of the calf. Weakness, dizziness and fatigue related to a moderate postural hypotension. This may be corrected by a decrease in dosage. Older age may cause slight ataxia, postural hypotension with faintness, especially. The drugs should be used with caution in patients who have a history of asthma or peptic ulcer. They may be given to a very old or partially paralyzed (paralytic) patient.

Dose	How Supplied	Dosage
Tolazoline HCl U.S.P. (Dibyl®)	25 mg tablet	Start with 1/2 tablet 4 times a day and gradually increase to 4 or 6 tablets daily
Phoxybenzamine HCl, NND (Dibyl®)	10 mg capsules	Start with 2 capsules daily, increase by 1 capsule every 4 days up to 4 or 6 capsules daily
Azepate Phosphate NND (Iliad®)	25 mg tablets	Start with 1 tablet 4 times a day for 1 week, 2 tablets 4 times a day for the 2nd week, then may increase to a maximum of 2 tablets 4 times a day
Nylidrin HCl NND (Aridin®)	6 mg tablet	Start with 1 tablet 4 times a day and increase gradually to 1 tablet 4 or 6 times daily

D Treatment of the Severe Stages of Peripheral Arterial De
compensation**1 Treatment of claudication**

- a Teach patient to walk slowly take short steps and to stop to rest before the pain of claudication is fully developed
- b Correct any ligamentous or arthritic disabilities stretching exercises salicylates

2 Treatment of rest pain

- a Have patient sleep with the head of his bed elevated 8-10 inches
- b Limit activities rigidly
- c If edema has developed, an oscillating bed or Buerger's exercises may be prescribed (see page 209)

3 Treatment of severe infection or incipient gangrene

- a Start antibiotics as soon as infection occurs (see page 514)
- b Keep extremity horizontal or lowered never elevated. The oscillating bed may be useful.
- c Keep the foot free of dressings
- d Room temperature must be comfortable (70°-80° F)
- e Support bed clothes by use of a cradle over affected limb or by a pillow under bedclothes at the foot of the bed
- f Drain purulent pockets thoroughly but gently. This may be accomplished by covering crusted lesion with a few layers of Vaseline® or Xeroform®; use for 24 hours then applying saline sponges at room temperature and changing frequently during the next 48 hours. Then dress the lesion with a bacitracin or bacitracin-neomycin ointment and a single layer of Xeroform® gauze for 2-3 days. Reinstitution of this treatment when necessary.

E Surgical Measures

- 1 Thromboendarterectomy is especially useful in the segmental or localized occlusion of major arteries
- 2 Sympathectomy if there is some evidence of abnormally increased vasomotor tone (see page 208)
- 3 Conservative amputation (toe or transmetatarsal) when the active hyperemia and elevation test shows a filling pressure in the small blood vessels of 35 cm or more (see page 508)
- 4 Classical supracondylar amputation if filling pressure in small blood vessels by reactive hyperemia test is less than 35 cm and thromboendarterectomy or sympathectomy is not indicated

VASCULAR SPASM**Treatment**

- A General Measures The same as for occlusive diseases. However tobacco in any form must be strictly prohibited.
- B Local Measures The same as for occlusive diseases especially if associated occlusion is present.
- C Measures aimed at prolonged or permanent relief of spasm
 - 1 Surgery Sympathectomy of the affected extremity is usually the treatment of choice.

Crit is for sympathectomy Best d terminated on clinical grounds assisted by vasodilator tests

(1) Clinical evidence of increased vasomotor tone This is evidenced by sweating cyanosis and constricted veins (absence of ever rubor and normal or slow blanching reaction on elevation)

(2) Sympathetic block imilla te t giv relief f pain and i termitt t claudication and better color to feet

b Co t aindications

(1) Mark d ubor on dep nde cy

() R p d blanching on lev tion

(3) At ophy of t ues

• Vasodilator d gs

a Chem al symp th et my The introduction of ganglio i bl cki g g nts has afford d a n w pproa h to the relief of b orm l aso st tion Many ha e b n tri d b t only a f w a usef l

(1) Adr glc blocking ag nt These dr gs ct at the n ve ndings in th va cul r muscle ells (n uro fi ct r s te) whi h s probably the most desi abl mode f act on They thus n t only blo k the sympa th ti v soco trict r stimu b t al o the vaso on str tor ffects f om ircu lating epin phrin and o pl eph ine Th s gro p of d ugs i el d s Tolazoline Hyd ochlorid U S P (P is ol n) ph o yb e za m (D b zyl e) Azap ti Phosphate N N D (lild) nd Nyl d Hyd ochl rid N N D (Ar lidin) (et ble) Th drugs a e eff tiv ally Th y a the most ef l drug unte a ting ab m l o p m

S de ffe t wh h a tro bles m but t s iou may b al co gest m s p cki s natio of ac lp W ak s d zin ss a d fatigue a e r lated to a m de te po rur l hvp t sion ths may b e r ted by a d e in do ag Ove dos g m y e s lti m p fo d post l hypoten o with fat tn ss y p Th drug should be used with c ut a y p t t who g ves a h at y of thm o p pt lce Th y m y be g e i t a enou ly o i t a terially (ly c ry)

D ug	H w bu ppl d	Dosage
T las lln HCl U S P (P i l n)	25 mg tabl ts	Start w th $\frac{1}{2}$ tabl t t d p and gr du ally e to 4 8 t bl t daily
Ph oxyb n rein HCl N N D (D ben yll)	10 mg ps l	St rt w th 2 p l d lly p and in a by l e y 4 d ys up to 4 6 c psules daily
Az p e tn Ph phat N N D (lild)	25 mg tabl t	St rt w th 1 t bl t t d p f r l w k 2 t bl t t id for the 2nd w k the m y e s to a m xim m f 2 tabl t q d
Nyl d HCl N N D (Ar lid)	5 mg tablet	St t w th 1 t bl t t d p and i a g d ally t l t bl et 4 6 t mes daily

- (2) *Veratrum viride* compounds Produce peripheral vasodilatation by depressing vasomotor centers in the hind brain. They do not block vasoconstricting effects from circulating epinephrine and norepinephrine. They are relatively ineffective in a patient with vascular spasm.
- (3) Tetraethylammonium ion and methonium compounds Block sympathetic and parasympathetic impulses at ganglionic synapse. They do not block vasoconstricting effects from epinephrine and norepinephrine and may potentiate vasoconstricting responses to epinephrine and norepinephrine.
- b Direct vasodilators (act directly on vascular muscle)
Nitrites, nicotinic acid and derivatives have not proved too successful in patients with abnormal vasoconstriction.

ACUTE ARTERIAL OCCLUSION (Acute Arterial Embolism code No 46 618)

Acute arterial occlusion is usually due to embolism. It occurs most commonly in patients with a circular fibrillation or myocardial infarction but may result from thrombosis of a vessel especially during periods of hypotension.

The onset is frequently sudden and associated with severe pain. Constitutional symptoms and shock are present if the artery is of large caliber. There is pulselessness of the distal artery and pallor and coldness of the extremity with numbness, tingling and muscle paresthesias. If treatment is not instituted the extremity or part may undergo gangrenous change.

Treatment

- A Surgical Immediate embolectomy is the treatment of choice.
- B General Measures To combat the thrombotic extension of the embolus and reflex vasoconstriction institute the following measures at once and continue postoperatively (reference if surgical treatment is not possible):
- 1 Morphine sulfate 10-15 mg ($\frac{1}{8}$ - $\frac{1}{4}$ gr) I.V. at once and repeat as needed subcutaneously.
 - 2 Anticoagulant therapy should be instituted at once to prevent thrombotic extension of the embolus. Give Heparin Sodium U.S.P. B.P. 2 cc (20 mg) (2000 unit) I.V. immediately. The effects of this heparin will usually have worn off by the time the patient has been transported to a hospital or prepared for operative treatment. The usual regimen of anticoagulant therapy is then started as soon as possible (see page 215).
 - 3 Procaine or xylocaine block of the sympathetic system to the affected extremity may be helpful. Repeat as necessary but use cautiously in the patient who has received anticoagulant therapy.
 - 4 Vasodilator and sedatives
 - a Papaverine Hydrochloride U.S.P. B.P. 30-60 mg ($\frac{1}{2}$ -1 gr) I.V. every 2-3 hours.
 - b Ethyl alcohol (as alcoholic beverage) orally in generous amount.

Ad energetic blocking agent (See page 211)

5. Dilating bed Useful in acute occlusions

C. Local Measures

1. Keep extremity horizontal or slightly depressed if an occluding bed is not available protect against pressure or trauma

2. Avoid use of heat or cold to the affected part

D. Treatment of Ischemic Neuritis May follow a useful routine

1. Cyanocobalamin U.S.P. (vitamin B₁₂) 1000 mcg hypodermically daily for 2 weeks has been advocated

2. Arteriotomy will give relief if Vitamin B₁₂ therapy does not help but establishment of circulation by thromboendarterectomy is preferred

DISEASES OF THE AORTA

AORTIC ANEURYSM

(Syphilitic code No 461 147 6)

(Arteriosclerotic code No 461 942 6)

Aortic aneurysm is a pulsating swelling which forms as the result of dilatation of the wall of an artery. The most important part of the aorta is the thoracic arch are most frequently affected. Thoracic aortic aneurysm is most commonly retroperitoneal.

The signs and symptoms vary with the location and size of the aneurysm. Most frequently they are due to local pressure on the surrounding structures. The most common symptom is pain which results from pressure on surrounding structures. Pressure on the trachea about the aorta can cause dyspnea and cough. Abdominal aortic aneurysm may produce back flanks or groin pain. Some aneurysms may be asymptomatic and may be discovered by physical examination or by x-ray of chest or abdomen.

Treatment

A. Spontaneous Malignant Treatment underlying syphilis if present (see page 440)

B. Surgical

1. Replace the weak wall by an autogenous vein or vein graft or homograft or synthetic material

2. Palliative Attempts to halt further dilatation by producing an intentional thrombosis of the wall of the aorta by external or internal means to arrest the dilatation to be used only in the low risk patient

DISSECTING ANEURYSM OF AORTA (code No 461 940 1)

Dissecting aneurysm is caused by the rupture of the intima and the formation of a false lumen in the presence of hypertension. It usually leads to a fatal aortic changes in the aorta.

It is manifested by a sudden onset of severe agonizing pain usually in or near the site of the rupture. The pain may radiate to the head, back, pelvis or legs. Shock follows rapidly and death usually occurs in a few hours or days. Diagnosis is usually made at autopsy because of the clinical similarity to myocardial infarction and acute arterial occlusion.

Treatment

Treatment is entirely symptomatic and similar to that of myocardial infarction (see page 185). Surgery has been used successfully in a few cases.

DISEASES OF THE VEINS

VENOUS THROMBOSIS (code No 48 619) THROMBOPHLEBITIS (code No 48 100 7)

A condition of uncertain etiology in which a thrombus forms in a small vein (usually the lower part of the leg) and grows by deposition of fibrin and cells, filling the larger veins of the leg (98% of cases). Inflammation of a localized area or much of the vein may be present. Early in the disease the chief danger lies in the detachment of all or part of the thrombus producing pulmonary infarction. Years later the chief danger lies in the development of the postphlebitic leg with edema, subcutaneous fibrosis and ulceration.

This condition is common in both medical and surgical patients. The present medical and surgical treatment methods are only probably of value but have now to be decided.

Diagnosis

Early diagnosis and immediate therapy are of importance to prevent pulmonary infarction.

A History Venous thrombosis tends to occur after abdominal or pelvic surgery, trauma, prolonged bed rest and in malignancy.
1 Pain in calf and behind knee as important early symptom.

2 Pleuritic pain, sudden onset with bloody sputum, suggestive of pulmonary infarction.

B Physical Examination May be negative.

1 Examination signs

a Diffuse increase in color of feet with levitation.

b Slight difference in temperature.

c Dilatation of superficial veins of leg.

2 Pain or tenderness on palpation of main venous channels of calf or foot. Do not palpate too vigorously.

3 Homans sign Limitation of motion of toes, dorsiflexion of foot.

4 Swelling of the calf. Usually late sign. May be diminished only by measurement and comparison with the opposite limb or by repeated measurements.

5 Examination of the chest in case of suspected pulmonary infarction may reveal signs of diminished breath sounds and crackles or pleuritic friction rub.

Treatment.

Anticoagulant Therapy. As soon as the diagnosis of venous thrombosis is made, anticoagulant therapy must be started at once. *Prothrombin level and Lee-White clotting time must be determined first.*

1. Heparin. Prolonged anticoagulant action of heparin may be obtained by the deposition of a highly concentrated solution of crystalline heparin into a relatively compact and avascular area of the subcutaneous fat. One injection daily appears to give a prolonged anti-coagulant action. A highly concentrated aqueous heparin (200 mg per cc) is injected slowly through a No. 25 needle into the subcutaneous fat 1 1/2 inches below the posterior iliac crest. Average doses are as follows:

100 lb patient 200 mg daily

150 lb patient 250 mg daily

175-200 lb patient 250-300 mg daily

Check Lee-White clotting time before starting treatment and just before the next dose. If the clotting time exceeds 18 min, defer the next dose until it falls below this level. Modify dosage as necessary.

At present the most general use of heparin is during the first stage of treatment from the first to the third days of anticoagulant therapy until the partial thrombin depressant becomes effective. The subcutaneous administration of heparin may be used alone without the addition of prothrombin depressants.

2. Prothrombin depressants. During the first stage of treatment (1-3 days) it is best to supplement the drug with heparin until prothrombin concentration has therapeutically levels (10-30%). Prothrombin levels should be determined daily and the next dose not given until the day level is known.
 - a. Bishydroxycoumarin (USP (Dicumarol®)). Usually takes 48-72 hours to reach effective therapeutic level and the amount of time to return to normal after discontinuing treatment. Initial dose is 200-300 mg on the first day, 100-200 mg on the second day. Maintenance dose varies from 25 to 150 mg daily.
 - b. Ethyl Biscoumatate (NND (Tromexan®)). Tromexan® is said to induce a more rapid fall in prothrombin concentration and a more rapid recovery of action on oral administration than Dicumarol®. Initial dose is 1500-1800 mg in 2 divided doses on the first day and 300-600 mg on the second day. Maintenance dose is 300-900 mg daily in divided doses. Heparin is usually only given for the first 24 hours because of the more rapid action of Tromexan®.
 - c. Phenindione (NND (Hedulin® (Dalon®))). Has the advantage of producing a definite action on total glycerophospholipids with Tromexan®. Initial dose 200-400 mg in 2 divided doses. Maintenance dose is 25-150 mg in divided doses. Vitamin K is apparently not effective in counteracting the effect of phenindione but this may not be important in view of the rapid return to normal prothrombin levels after withdrawal of the drug.
 - d. Warfarin (Sodium NND (Coumadin®)). Is a hydroxy coumarin derivative which may be administered orally or parenterally. The dosage is the same irrespective of the route of

administration. The initial dose is 75 mg orally I V or I M. The maintenance dose is 5-10 mg daily. Induction of prothrombin depression is rapid but the return to normal may be prolonged up to 5 days.

- 3 Duration of therapy varies with each case. For most patients this is about 10-14 days. Continue the therapy for about 7 days after there is no further fever or pain.
- 4 Treatment of bleeding and over dosage. The principal danger from anticoagulant therapy is abnormal bleeding.
 - a Bleeding due to excess heparin. Discontinuing the therapy will usually terminate bleeding in about 1-3 hours only if I V heparin has been used. If immediate cessation is necessary, slow I V injection of protamine sulfate 40-50 mg will neutralize the effect of 50 mg of heparin.
 - b Bleeding due to excess prothrombin depression is more difficult to control. For the prothrombin level rises slowly after therapy is discontinued. The rise is more rapid when Tromexan® or phenindione has been employed.

(1) Severe bleeding

(a) Stop the drug and do not use again.

(b) Fresh blood (clotted) if available immediately.

(c) Phytonadione U S P (Mephyton®) 50-200 mg I V slowly (at rate not over 10 mg/min) but by syringe or added to venous clots of dextrose and/or saline) and repeat every 6 hours as necessary. The natural more rapidly than synthetic vitamin K like product (as menadiol) is better.

(d) Mephyton Sodium Bitartrate U S P. Give 50-100 mg I V Stat and repeat 2-3 times the first day.

(2) Mild bleeding (mild nose bleed, hematuria, urinary oozing)

(a) Stop drug, rest at low dosage, stop prothrombin time rises to 20-30".

(b) Phytonadione U S P (Mephyton®) 5 mg (or 10 mg) 3-5 times daily. Usually 5-30 mg daily. If bleeding is not controlled or becomes more severe use I V as above.

c Overdosage of depression with bleeding. If the prothrombin level drops below 10% and does not rise 2 days after discontinuing administration, give Mephyton U S P (Mephyton®) 5-30 mg orally. When prothrombin rises the drugs may be given again.

B Venous Ligature

1 If anticoagulant therapy is contraindicated, venous ligation is recommended for a variety of conditions. It is a permanent therapy and is contraindicated. The same is true with pulmonary embolism. The procedure of ligation is best in cases of renal or hepatic disease and in cases of peripheral C N S injury.

2 Active thrombus or embolism form. Venous ligation holds the perfused distal part of the peripheral part of the thrombus. If embolism occurs to occur while under anticoagulant therapy or if septal phlebitis is present.

C Garter or Pile Treatment in bed with foot elevated 4-6 inches. An elastic bandage is applied snugly from foot above knee to mid thigh to keep venous outflow. Do not obstruct arterial circulation. Check pulses. R wrap every 6 hours.

- 1 Exercise As soon as treatment is started allow for the minimum and exercise in bed (see below). If leg is in a cast put the foot on a stool by turning and relaxing muscles in bed.
- 2 Ambulation As soon as the acute pain subsides (or if no pain is present as soon as therapy is instituted) the patient must be made ambulatory (unless other systemic conditions prevent this). During this time an elastic bandage should be worn. The time out of bed and walking is increased very gradually. The elastic bandage should be worn for about 3 weeks after full ambulation has been achieved.

Prophylaxis

A Early Ambulation and Exercise

- 1 Early ambulation Prolonged bed rest or inactivity should be avoided especially in elderly patients. Have patient up and about as soon as possible after operation or a useful illness. Walking a few steps is preferable to sitting for half an hour in a chair.
- 2 Bed exercises If bedrest is necessary pass the legs or active bed exercises should be instituted as soon as possible and should be continued as long as patient must remain in bed. The contraction of the calf muscles is sufficient to move the ankle and hip repeat 5-10 times every hour while awake.
- 3 Movement in bed With patients at bedrest keep bedclothes loose and patient moving slowly.
- 4 Elevation and massage Elevation of the foot of the bed 4-6 inches and wrapping legs from the toe to just below the knees with a bandage will usually promote venous return.

B Routine Prophylactic Use of Anticoagulants In elderly patients who cannot be fully mobilized the above regimen should be of value (see also the chapter on page 215) but in general the routine prophylactic use of anticoagulants is not advised.

POSTPHLEBOTIC EDEMA AND ULCERATION

Almost all patients with venous thrombophlebitis of a lower extremity develop the postphlebotic syndrome eventually. In developing the syndrome a predisposing disability the haemostatic status of the syndrome is a factor. (1) reanastomosis of the deep veins of the lower extremity with incompetent deep valves and edema or superficial varices or more frequently both (2) subcutaneous inflammation to the thigh (usually in the medial aspect of the lower third of the thigh) with the usual subcutaneous fibrosis atrophy pigmentation and sclerosis of the overlying skin and (3) ulceration.

Diagnosis

A History The history may not be a history of thrombophlebitis but there is almost always a history of major surgery or fracture of the lower extremity or prolonged bed rest for medical reasons.

B Symptoms and Signs

- 1 Edema and changes of the leg after prolonged standing going sitting is a frequent symptom. It may be relieved or improved by elevation of the extremity or by walking. Congestion and discoloration of the skin may be present.
- 2 Increased pigmentation of the superficial veins.

212 Postphlebitic Syndrome

- 3 Episodes of inflammation of the medial aspect of the lower third of the leg
- 4 Dry scaling eczema of the lower medial part of the leg
- 5 Varices of the superficial saphenous system and incompetence of the deep valves usually can be demonstrated
- 6 Brownish red pigmentation of the skin just above the medial malleolus or less often just below the lateral malleolus
- 7 Subcutaneous fibrosis with retraction and shrinking of the areas may be very painful and disabling. Fibrosis is usually just above the medial malleolus but may girdle the leg
- 8 Ulceration in areas of subcutaneous fibrosis

Treatment

The accumulation of protein rich edema fluid is the main cause of the more serious manifestations of the postphlebitic syndrome. If edema is conscientiously avoided these may be prevented.

A Control Edema

- 1 Sleep every night with foot of bed on 4 inch blocks
- 2 Wear well fitted (made to order NOT stock supply) heavy duty elastic stocking below the knee with fitted heel
- 3 Three to four 20 minute rest periods during the day with feet elevated 6-8 inches above heart level

B Control Infection Control of dermatophytosis and any bony coses is essential. Castellani's dye to toes and nails once or twice a week is probably the best control measure.

C Elevation and Ulceration Once these signs appear elastic support is not adequate. A carefully fitted semi rigid boot of the Unna's paste boot type will heal most ulcers in 4 months. Boots may be applied with tape and sheet cotton Vaseline[®] or Gaultex[®]. The patient continues his usual activities. The boots should be applied with firm venous pressure over the leg without irregularities which may cause further damage to the skin. They must be changed every 1 to 2 weeks depending on drainage but once the ulcer is healed or drainage is minimal it can be left on as long as 4 to 6 weeks.

Viscopaste[®] bandage is a Unna's paste type bandage 3 1/2 inches wide impregnated with gelatin and zinc oxide. The Gaultex[®] bandage is a 3 inch bandage impregnated with a self adhering compound which is surprisingly non allergic.

The bandage should extend from the base of the toes include the heel and continue to a point immediately below the bend of the knee. A thin layer of cotton sheeting or gauze is used to pad the Achilles tendon and the dorsum of the foot. An extra layer of cotton or gauze and sometime a bobby sponge 1/2 inch thick is placed over the ulcer. Special attention especially antibiotic preparations are not necessary. The bandage is started with one horizontal turn around the foot and when completed it is carried obliquely over the heel and then back around the foot. After the heel has been adequately covered the bandage is carried up the leg. No attempt should be made to apply it as a continuous spiral. It should be allowed to follow its own course without pleating and should be retied frequently if necessary so that bandaging may be recommenced to build up a uniform thickness of paste or bandage which should be molded carefully to conform evenly to the shape of the limb.

Chapter 9

DISEASES OF THE BLOOD AND LYMPHATIC SYSTEMS

ANEMIAS

HYPOCHROMIC ANEMIA (code No 501.735) (Normocytic and Microcytic)

Hypochromic anemias (I w color index and M C H of less than 27 yr) include anemias due to nutritional deficiencies, infections and to iron and chronic blood loss. Women require about 4 times as much iron as men up to the menopause.

Pathogenesis

A Chronic blood loss

B Inadequate intake of iron (nutritional anemia)

C Defective absorption of iron in the gastrointestinal tract (e.g. hypochromic anemia of infection). Factors influencing absorption of iron include the following:

1. Acorbic acid facilitates the absorption of iron hydrochloride and does not

2. Depletion of body iron increases the absorption of iron

3. Valence of iron ingested (Fe^{++} is easily absorbed, Fe^{+++} is not)

4. Infection usually decreases absorption of iron

5. Potassium deficiency also decreases absorption of iron

D Idiopathic mechanisms

E Pregnancy

Treatment

A Specific Measures

1. Correct the factors causing or contributing to anemia (e.g. hemorrhage, infection, diarrhea, vomiting, intestinal parasites, abdominal distention, diaphragmatic hernia, gastric polyps, and tumors)







2. Iron is the chief factor in the type of anemia. It is best given immediately after meals

a. Orally, per se and dosage (Physiological need is 15 mg/day; maximum absorption is considered to be about 100 mg/day)

(1) Ferrous Sulfate U.S.P. B.P. 0.2-0.3 Gm (3-5 gr) tid, p

(2) Ferrous Sulfate Syrup U.S.P. 0.12 Gm 2 gr per tsp) 4-8 cc (1-2 dr) bid or tid p.c.

SIZE COLOR RELATIONSHIPS OF RED BLOOD CELLS IN THE VARIOUS ANEMIAS

SIZE	COLOR		
	HYPOCHROMIC MCH < 21 γγ CI < 0.9	NORMOCHROMIC MCH 27.32 γγ CI 0.911	HYPERCHROMIC MCH > 30.77 CI > 1.1
MICROCYTIC MCV < 78 cμ VI < 0.9	Anemias due to faulty GI absorption of iron Iron deficiency anemias Anemias due to infections Anemias due to chemical and physical toxins Splenic anemias Erythroblastopenia	 Uncommon	 Uncommon
NORMOCYTIC MCV 80-94 cμ VI 0.911	Anemias due to blood loss Hemolytic anemias Erythroblastic anemias Myelophthisic anemia	 As for hypochromic microcytic and normocytic anemias Aplastic anemias	 Uncommon
MACROCYTIC MCV > 95 cμ VI > 1.1	Macrocytic anemias complicated by iron deficiency or blood loss or other of the above factors	 As for hyperchromic macrocytic anemias	 Pericious anemia Tropical anemias Anemia of pruritus Fish tapeworm anemia Macrocytic anemias associated with Idiopathic atrophic gastritis Chronic liver disease Gastric carcinoma (rare) Faulty GI absorption Lukemia Pregnancy Infancy

- (3) Ferrous Carbonate Pills N.F. P.H.O.I. Car
bonate B.P. 0.510 Gm (7 1/2 15 g) tid p
(4) Ferrous ammonium sulfate solution (50%) 4 cc tid p
(5) Ferrous Glucose USP 0.3 Gm (5 g) tid p
b. Parenteral Iron Indication: Intolerance to oral iron
refractory to oral iron (poor absorption) gastric
intolerance, especially use of alkaline phosphate
of depleted iron stores where oral iron fails ade-
quately to oral iron. The following forms of parenteral
iron hold the edge only: iron dextran complex or
iron dextran complex. I will list this method by the following
formula: 0.55 weight (lbs) x (normal Hgb % - patient
Hgb %) mg of metallic iron (total dose) needed for
(1) Sodium iron oxalate I.V. concentration 2%
metallic iron (0 mg per cc) Give 50 mg (3/4 g) *has*
Stat and then 100 150 mg (1 1/2 1/2 g) I.V. daily *#1*
until the total dose has been given
(2) Iron Dextran Complex N.N.D. (Imferon) for I.M.
concentration 5% in tall iron (50 mg per cc) Give
50 mg (3/4 gr) Stat and then 100 150 mg I.M. daily
or very old day until the total dose has been given
life time plus with a 2 inch needle into the posterior
quadrant of the buttock using the Z-track technique (pulling
skin to needle before injecting) to prevent
leakage of the old iron and discoloration of the skin

3. Adjunct therapy

- a As bc Acd USP G ea ng J c or
orb c d t blet 30 50 mg p d y fo child en
100 150 mg per d y f ad lt
- b D lut d Hydr hl Acd NF (10%) 2 4 {1/2 I
dr j t i d in gl ss of w ter w th m e l s p p d thro gh
a gl s f b e tube fr qu ntly has b e p s i b d for
pat nts w th chl hydr a but r c e t v i n n d s t s
th t d l t hvd chlo ac d do not f e l t t th ab
o p t i o n of in th g t r o n t e s t a l t a t I f p c i b e d
pat n t m u t h u h t t w th s o d u m b r b o n a t a f t e
m i s t n t a l c d i e f t o n t e t e t h

B G 1 M 8

1. Diet High in high protein on high vitamin
at least 70 Gm protein/day for average adult Food
high in on include liver other organ meats fresh
meat yeast eggs egg yolk especially vegetable
protein potato pumpkin A2440 low diet contain
ing 70 Gm protein 115 Gm fat and 230 Gm carbohydrate
with protein 20 mg of iron
2. Vitamins Vitamin deficiencies usually multiple and a
associated with the nutritional deficiencies Make a
full survey for nutritional status before diagnosis exp
iv vitamin preparation at which are of indicated
a. Iodine Vitamin D deficiency diet too diet
associated with anemia ludibb pellagra scurvy
achyophthalmism diet itami Adf cey
b. Deficiency causes a most often multiple so that
ly divisible to administer in its vitamin preparation
Specific preparation Usolyfrase if vitamin

deficiency (see page 59)

- 3 Whole blood transfusions preferably of fresh blood are used when there is need for rapid restoration of hemoglobin. this need is more urgent when hemoglobin is less than 8 Gm per 100 cc (30%) (see page 247)
 - a Acute hemorrhage when blood loss is greater than 500 cc
 - b Chronic hypochromic anemia when
 - (1) Need for correction of anemia is urgent (e.g. surgery and acute sepsis)
 - (2) Failure to respond to anti anemic measures. Re evaluate and consider other causes for the anemia (e.g. blood dyscrasias and serious constitutional diseases)
- 4 Red cell mass (sludge) transfusions are used to restore hemoglobin and red cells without increasing plasma volume and perhaps reducing risk of serum reactions
- 5 Thyroid May be indicated if anemia is associated with frank hypothyroidism or myxedema (see page 368)

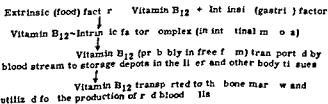
MACROCYTIC ANEMIAS

PERNICIOUS ANEMIA (code No 501.702)

Pernicious anemia (P.A.) is a chronic and if untreated progressive macrocytic anemia. The primary defect is the failure of the gastric mucosa to produce a substance (intrinsic or gastric factor) that is essential for the absorption of the vitamin B₁₂ in certain foods.

In the absence of intrinsic factor inadequate absorption of vitamin B₁₂ occurs and a deficiency of this vitamin develops. Vitamin B₁₂ is essential for normal erythropoiesis. When deficient pathologic red cell formation (megaloblastic regeneration) occurs. There is no primary deficiency of folic acid in P.A. The disease responds specifically to the parenteral administration of vitamin B₁₂ or of extracts of liver containing vitamin B₁₂ or to the oral administration of live or freeze preparations containing intrinsic factor and vitamin B₁₂.

The relationship of dietary and gastric factors to normal blood formation (modification of Castle theory) may be outlined as follows:



Diagnosis

A Symptoms

- 1 Anemia Weakness dyspnea and palpitation
- 2 Gastric intolerance Anorexia diarrhea and dyspepsia
- 3 CNS Numbness and tingling of extremities and sphincter incontinence

B Signs Pallor, icteric tint, tachycardia, glossitis, mild hepatomegaly and plomalgia, diminution of vibratory sense and ataxia

C Laboratory

- 1 Hematime achlorhydria
- 2 Macrocytic anemia
 - a MCV 95-160 μ
 - b MCHC > 32-36% (normal)
 - c MCH 33-38 $\gamma\gamma$ (normal)
 - d Orthochromatic megablasts (normoblasts) presentAnisocytosis, poikilocytosis and polychromatophilia
- 3 Bone marrow changes
 - a Reticular increase in size and soft
 - b Large numbers of megablasts are present

D Therapeutic Response to Vitamin B₁₂ Extract of Liver and Lactobacillus

- 1 Disappearance of symptoms and signs
- 2 Reticulocyte response (normal count is less than 1%)
- 3 Improvement of anemia Occurs about 1 week after beginning adequate parenteral or oral administration of vitamin B₁₂ therapy

Treatment

Treatment should be based upon an accurate diagnosis. Different minimum other conditions characterized by macrocytic anemia.

A Specific Measures Parenteral therapy is obligatory for those patients who cannot or will not accept parenteral treatment. Neither crude liver extract nor folate should be administered to P.A. patients.

The parenteral administration of refined extract of liver or of vitamin B₁₂ uniformly followed by optimal clinical and hematological responses. Following such therapy the P.A. patient in remission will undergo an increase in circulating reticulocytes (peaking peak in 6-10 days) and a return of erythrocyte and hemoglobin values to normal in 3-5 weeks.

1 Parenteral dosage and indications

a) Full compliance with P.A. in remission

- (1) Initial IM injection 20-30 mg of crystalline vitamin B₁₂ (Cyanocobalamin U.S.P.) vitamin B₁₂ once a day for first 3-5 days, then 10-20 micrograms of vitamin B₁₂ per day given the equivalent of 20-30 microgram of vitamin B₁₂. Vitamin B₁₂ concentrations may cause allergic reactions.
- (2) Subsequent injections
 - a) Patient in remission Give 15-20 micrograms vitamin B₁₂ equivalent every third fourth day

until blood values return to normal

(b) Mild relapse 15-20 micrograms weekly usually is adequate

- b For P A complicated by degeneration of the spinal cord Doses in excess of the amounts needed for uncomplicated P A may be required. The degree of reversibility of neurologic manifestations is inversely proportional to the duration of symptoms. Improvement frequently is marked in patients with symptoms of 6 months duration or less, less pronounced in patients with symptoms of 6-12 months duration, and negligible in patients with symptoms of more than 1 year's duration. It is advisable to treat all patients intensively for at least 6 months and preferably for 1-2 years.

Physical therapy including coordination exercises is an important adjunct to the specific therapy of P A complicated by spinal cord degeneration.

(1) Initial I.M. injection 30-60 micrograms of vitamin B₁₂ or refined liver extract

(2) Subsequent injections 20-30 micrograms 2 or 3 times weekly for 6 months or more or until optimal neurologic improvement has been demonstrated. If optimal neurologic improvement has not occurred at the end of 6 months continue with 20-30 micrograms once a week.

- c Maintenance therapy. The nutritional requirement for vitamin B₁₂ in normal individuals is 1.0 microgram or slightly less each day. This amount administered to patients with P A in whom blood values have been returned to normal and optimal neurologic recovery has been observed will provide satisfactory control in most instances. 15 micrograms of vitamin B₁₂ or refined liver extract I.M. once every 2 weeks should be adequate. Larger doses may be necessary during periods of increased stress (infection, prolonged debility, or other illness).

The patient must be instructed as to the need for adequate and regular supply for the remainder of his life. The serious risks of neglect should be emphasized.

Oral preparations. The response to the oral administration of powdered high gastric stomach preparations and tablets of vitamin B₁₂ (even in high dosage) usually is slower, less uniform, and often suboptimal when compared with the response to parenteral therapy. It is possible that the oral administration of intrinsic factor-vitamin B₁₂ complex substances or the inhalation of powdered vitamin B₁₂ following intranasal dosing will prove to be satisfactory methods of treatment, but the reliability of these methods has not been proved as yet.

- a Powdered Stomach U.S.P. (Ventriculi [®])
- b Lifer With Stomach U.S.P. (Ext. [®])
Tablets of vitamin B₁₂
- d Powdered vitamin B₁₂
- e Combination vitamin B₁₂ With Intrinsic Factor C.N.C. [®] (dried from various sources)

B General Measures Periodic clinical and blood examinations should provide the basis for administration and dosage of liver iron and adjuvant drugs

- 1 Rest Patients with profound anemia should be at bed rest Hospital care may be necessary for patients with neural involvement (ataxia and sphincter disturbances)
- 2 Diet A diet adequate in calories minerals and vitamins does not need to be supplemented with extra quantities of dietary liver unless for some reason the patient is not receiving parenteral liver therapy
- 3 Iron Ferrous salts may be given as an adjuvant to liver therapy when the iron content of the red cells is low ($\text{MCHC} < 32\%$ or low color index) Elderly patients usually require more iron (see page 219)
- 4 Diluted Hydrochloric Acid N.F. (10%) 24 cc ($\frac{1}{2}$ l dr) diluted in a glass of water taken through a glass straw with meals may be given to patients who have diarrhea as a complication of the achlorhydria Patients must brush teeth with sodium bicarbonate immediately after meals to neutralize hydrochloric acid and prevent erosion of teeth
- 5 Thyroid may be used in patients who fail to respond due to associated hypothyroidism
- 6 Measures to improve liver function in P.A. patients with associated hepatic damage have been suggested in an attempt to aid utilization and storage of the extrinsic factor (see page 220)

MACROCYTIC ANEMIA OF PREGNANCY (code No 501.701) (Pernicious Anemia of Pregnancy)

A hyperchromic macrocytic anemia characterized by megaloblastosis in the bone marrow usually occurring at end of the second or during the third trimester of pregnancy

Treatment

This anemia responds specifically to the oral or parenteral administration of folic acid or to crude liver extract (which contains folic acid) It does not respond to the parenteral administration of vitamin B₁₂ or refined liver extract (which contains virtually no folic acid)

A Specific Measures After delivery treatment with folic acid (or crude liver extract) may be discontinued since relapse does not occur

- 1 Folic Acid U.S.P. 10-15 mg ($\frac{1}{10}$ - $\frac{1}{4}$ gr) orally daily
- 2 Liver Injection Code L.S.P. 4 cc (1 dr) I.M. daily
- 3 Vitamin B₁₂ and refined liver extracts are of no value

B General Measures

- 1 Give in a standard units of animal protein even if diet is already adequate begin early and continue throughout pregnancy
- 2 If hypochromia occurs ferrous salts should be administered (see page 219)

SPRUE

(Anemia of Sprue code No 501 703)

Sprue is a chronic disease of undetermined cause (probably due to nutritional deficiency) characterized by sore mouth glossitis indigestion and recurrent diarrhea with steatorrhea it results in anemia asthenia emaciation and even death. The anemia may be microcytic hypochromic normocytic hypochromic or macrocytic hyperchromic (megaloblastic).

Treatment

A Specific Measures

- 1 For hypochromic anemia Oral or intravenous administration of iron (see page 219)
- 2 For macrocytic hyperchromic anemia (with megaloblastosis)
 - a Corticotropin (ACTH) or one of the steroids (see p 424) are important advances in the treatment of this form of anemia in non-tropical sprue. Improvement in the anemia is thought to be the result of increased absorption from the gastrointestinal tract of nutrients in food including hemopoietic factors (vitamin B₁₂ folic acid etc.)
 - b Alternative therapy If megaloblastic anemia of sprue fails to respond to steroids or corticotropin (ACTH) therapy give one of the following
 - (1) Vitamin B₁₂ U S P 15-30 micrograms I M 1-2 times per week until remission is obtained and then 10-15 micrograms I M every 1-2 weeks
 - (2) Folic Acid U S P 10-15 mg ($\frac{1}{6}$ - $\frac{1}{4}$ gr) daily orally or preferably I M
 - (3) Liver Injection Crude U S P 4 cc (1 dr) I M daily

B General Measures

- 1 High alcohol high protein low fat high vitamin diet
- 2 Plasma and blood transfusions nitroglycerin for severe hypoproteinemia and anemia
- 3 Corticotropin (ACTH) or the steroids may be used in the hypochromic form in doses given for correction of the megaloblastic anemia and especially (see page 423). These substances increase the absorption of nitrogen fat and other nutrients from the gastrointestinal tract
- 4 If hypoproteinemia persists Menadione Sodium Bisulfite U S P 10 mg ($\frac{1}{6}$ g) I M or I V Stat followed by 5 mg ($\frac{1}{12}$ gr) orally b.i.d. or vitamin K₁ tablets (Mephyton®) 5-15 mg ($\frac{1}{12}$ - $\frac{1}{4}$ g) or more orally daily
- 5 Calcium Chloride Phosphate or Glutamate U S P 2 Gm (30 gr) daily i.d. and vitamin D 5000-20 000 units if hypocalcemia or tetany exist
- 6 Vitamin replacement by mouth

OTHER MACROCYTIC ANEMIAS

This group includes (1) nutritional macrocytic anemia (2) megaloblastic anemia of infancy and (3) megaloblastic anemias secondary to disease of or operative procedures on the gastrointestinal tract

Treatment

- A Specific Measures Give folic acid crude liver extract or vitamin B₁₂ as for spleen (see page 226)
- B General Measures Provide an adequate high protein high vitamin diet

APLASTIC ANEMIA (code No 501 900 0)

An acute or sometimes chronic disease of the hemopoietic system characterized by an altered production of red blood cells resulting from a depression or exhaustion of the bone marrow. The condition may be secondary to known marrow poisoning but also occurs in the primary or idiopathic form.

Diagnosis

- A History of exposure to marrow toxins (e.g. hemolysins, certain drugs and irradiation) is often obtained. Aplastic anemia is a persistent progressive anemia which like certain other anemias fails to respond to liver iron or diet therapy. Other causes of anemia cannot be demonstrated. Bleeding tendency is common.
- B Laboratory Finding
- 1 Anemia is usually normochromic normocytic
 - 2 Bone marrow often (not invariably) shows aplasia with fatty and fibrous replacement
 - 3 Leukopenia and thrombocytopenia are usually marked

Treatment

- A Specific Measures None are known
- B General Measures
- 1 Transfusion. Repeated transfusion with a fully typed and cross matched whole blood may prolong life for variable periods. Rarely patients who quit and cease all gain number of blood transfusions may go into spontaneous remission for a while.
 - 2 Discontinue all unnecessary medication.
 - 3 Remove patient from exposure to suspected toxins.
 - 4 Diet. Provide diet with adequate vitamins and minerals.
 - 5 If alive and if antithrombotic therapy should be given an adequate folic acid is of little value in true aplastic anemia.

HEMOLYTIC ANEMIAS

Classification (Modified after Dameshek)

- A Hereditary Defect of red blood cells shows an apparent morphological defect with hereditary susceptibility to destruction of hemolytic reactions
- 1 Spherocytosis (familial) anemia (spherocytosis) (congenital hemolytic anemia) (code No 513 092)
 - 2 Thalassemia (Mediterranean Cooley) anemia (familial erythroblastic anemia) (code No 501 997)
 - 3 Sickle cell anemia (Negro) (code No 513 94)
 - 4 Other disorders with abnormal hemoglobins

- B Acquired Defect (Acquired Hemolytic Icterus)** (code No 513 911 0) Red blood cells are originally morphologically normal. Etiology includes the following:
- | | |
|--------------------------------------|-------------------------------------------------|
| 1 Infections bacterial and protozoal | 4 Immune hemolysins |
| 2 Toxins venoms drugs and chemicals | 5 Agglutinins |
| 3 Physical agents | 6 Abnormal splenic mechanisms hypersplenism etc |
| | 7 Certain ovarian cysts |
- C Unknown Defect** (code No 513 900 9)

Diagnosis**A History**

- 1 Familial or racial hemolytic tendencies (hereditary)
- 2 Exposure to infections toxins agglutinins (acquired)
- 3 Symptoms of anemia (weakness dizziness palpitation and dyspnea) and hemolysis (fever chills abdominal pain and muscle cramps)
- 4 The acute hemolytic crisis is characterized by sudden onset of fever anemia icterus splenomegaly with tenderness and shock

B Physical Examination Pallor icterus tachycardia and fever may be present in all types. Splenomegaly and hepatomegaly occur in the acquired and hereditary types.

C Laboratory Findings

- 1 Increased blood destruction gives rise to
 - a Normocytic anemia
 - b Bilirubinemia
 - c Hemoglobinuria (see table on page 230)
 - d Increased urinary urobilinogen (urine dark)
- 2 Morphologic red blood cell defects Spherocytes target cells or sickle cells (see laboratory manual)
- 3 Altered red blood cell fragility (always present in hereditary types)
- 4 Increased blood formation is evidenced by bone marrow hyperplasia reticulocytosis and presence of immature erythrocytes
- 5 Leukopenia sometimes present in the acquired form
- 6 Detection and identification of abnormal hemoglobin which give rise to hereditary trait and anemia. This requires specialized chemical and electrophoretic techniques

Treatment of Acute Hemolytic Anemia (Hemolytic Crisis)

Patient must be hospitalized to Hospitalize if possible

A Severe Form

- 1 Treat shock (see page 28) and anemia. Careful observation of clinical progress is essential.
- 2 Whole blood transfusion. Blood must be carefully typed (major group and Rh type) and cross matched both at room and body temperatures. Severe reactions may occur even with a careful cross matching. Sturge recommends a cautious preliminary administration of 50 cc of suitable blood followed by an observation period of 1 hour. If no reaction occurs the remainder of the blood may be given over a 2 to 3 hour period.
- 3 Plasma. If patient cannot tolerate whole blood transfusions (because of hemolysis of injected red cells) give plasma transfusions when necessary to combat shock.

- 4 Specific us s should be treated with known
- Infectious Employ specific ant infective and supportive measures (see pages 496 to 514)
 - Discontinue drugs or remove from contact with poisons or other hemolysins
- 5 Corticotropin (ACTH) and the steroids may produce striking remissions of the hemolytic reaction and at least temporarily tide the patient over until such time that other more specific measures can be safely instituted. For dosage of ACTH and cortisone see page 424.
- 6 Splenectomy After shock and fever have subsided and patient's general physical status has improved sufficiently consider for early splenectomy. If hemolytic shock is progressive despite vigorous supportive measures (up to 3 to 4 500 c.c. transfusions) emergency splenectomy may be indicated. When the cause is unknown and reaction has been severe consider for splenectomy after patient has recovered from the hemolytic crisis. Splenectomy is not generally as successful in the acquired form as it is in the familial type and is generally without benefit in sickle cell and familial erythroblastic anemias.
- B Mild Form If the hemolytic reaction is mild only treatment with antinfective agent and corticotropin or cortisone may be necessary. In the familial type even though the patient is asymptomatic splenectomy may be advisable.

Treatment of Chronic Ph

- A Instruct patient to avoid strenuous exercise, infection, exposure to temperature extremes and ingestion of certain drugs or toxins.
- B Splenectomy If patient fails to improve on conservative therapy consider for splenectomy (see above). When abnormal antibodies (iso and auto antibodies) are present the indication is less often amenable to splenectomy.
- ✓ C Cobalt chloride, 200 mg orally daily has been employed in the early phases of sickle cell and Cooly anemia. A definite improvement is usually in effect with this same hard fast altering thyroid function.

HEMOGLOBINURIAS

Diagnosis (See table on the following page)

Treatment

- Specific Measures Remove or treat causative factor
 - Proximal old hemoglobinuria. Treatment of syphilis page 440
 - Favism. Prophylactic treatment of favism
- Symptomatic and Supportive Measures
 - Hemolytic symptoms
 - Tetracycline hemolytic reaction (see page 228)
 - Treat fever, chills and muscular aches and pain symptomatically
 - Anemic symptoms. Test acid to type and severity

DIAGNOSIS OF HEMOGLOBINURIAS

Disease	Precipitated By	Positive Laboratory Tests
Paroxysmal cold hemoglobinuria (code No 510 500)	Chilling or cold	Blood test for syphilis Don'th Landsteiner test
Paroxysmal nocturnal hemoglobinuria (code No 510 500)	?	Acid hemolysis test Hemosiderinuria test
Favism (code No 010 3761)	Ingestion of fava beans	None
March hemoglobinuria (code No 510 500)	Exercise	None

Prophylaxis

- A Paroxysmal Cold Hemoglobinuria Protect against chilling or cold
- B March Hemoglobinuria Avoid strenuous exercise

POLYCYTHEMIA VERA (ERYTHREMIA) (code No 501 79)

A chronic disease of the hemopoietic system of unknown etiology characterized by overactivity (erythroblastic) of the bone marrow with resultant overproduction of red cells and hemoglobin. It is manifested by a reddish purple hue to the skin, increased blood volume, capillary engorgement, hemorrhages, venous thrombosis, arterial hypertension, hepatomegaly and splenomegaly and symptoms referable to multiple organ systems. It is to be differentiated from the polycythemia that may occur secondarily to known physiological states which also cause increased bone marrow activity.

Treatment

- A Definitive Measures To reduce the total red blood cell volume
- 1 Venesection (phlebotomy)
 - a Utilize careful blood hematocrit determination in following efficacy of treatment
 - b Remove 500 cc of blood daily until the blood hematocrit reaches a normal level. Subsequently 500 cc phlebotomy every 2-3 months may be sufficient to control mild cases
 - 2 Irradiation Inhibition of red cell formation
 - a Radioactive phosphorus (P³²) This is the most effective anti polycythemic agent available at present. Its use is restricted to institution equipped to handle radioactive material. It is indicated in patient in which the polycythemia cannot be controlled readily by venesection alone and especially in patients with a history of thrombotic or thrombophlebotic episode. 4-6 milluri of P³² (as phosphate salt) in 2-6 cc of isotonic (1/2-1 1/2 dr) sodium phosphate solution is given I.V. If the polycythemia is not controlled following a single I.V. injection subsequent injections of 3-6 millicuries are given at intervals

of 2 months until the disease is brought under control

- b X ray irradiation Whole body or spray irradiation may be of benefit when given in repeated dosages. Irradiation of the long bones has proved to be less satisfactory than whole body irradiation in controlling the disease

3 Anti polythemic drugs

Phenylhydrazine hydrochloride or acetylphenylhydrazine. Follow patient carefully clinically and with blood studies during and after therapy. These compounds are most safely used if they are administered as maintenance therapy after the hematocrit has been reduced to normal by repeated venesections. Give 0.1 to 3 Gm (1½ to 5 gr) by mouth weekly as a maintenance dose. The use of phenylhydrazine to lower an elevated erythrocyte count, omitting the use of venesections to establish normal red blood cell and hematocrit levels is a hazardous procedure. Anorexia, nausea and vomiting are the principal disadvantages of phenylhydrazine therapy.

- b Triethylethylamine (TEM) has been employed but experience has been limited

B General Measures

- 1 Provide symptomatic relief as needed
- 2 Diet. The diet should be adequate and nutritious. There is no rationale for starvation diet or diet excluding no meat amounts of blood building foods
- 3 Inform patient regarding the nature of his disease

C Treatment of Complications

Varies with the status of the pancytopenia and the relation of complications to therapy as well as with the nature and site of the complication. Thrombosis and hemorrhage are common complications.

ACUTE AGRANULOCYTOSIS (code No 502.7911) (Agranulocytic Angina)

An acute and fulminant disease usually fatal in adults characterized by extreme granulocytopenia which is followed by a fulminating epistaxis associated with ulceration of skin and mucous membranes. It is known to be caused by certain drugs and chemical but is sometimes of unknown origin.

D Diagnosis

A History of Medication With Certain Drugs

Sulfonamides, pyriminyls, thiazides, phenothiazines, phenolphthalein, thiouracil and related compounds, gold salts, other heavy metals, estrogenic steroids, Tridione, butabarbital, Thioridazine, barbiturates (? rare), aspirin (? rare).

B Physical Examination

Sudden onset of pain and fever in inflammation and ulceration of mucous membranes of throat and locally of other areas of the skin and gingival and nail.

C Laboratory Findings

Leukopenia and granulocytopenia

232 Agranulocytosis

- a WBC usually $< 2500/\text{cu mm}$
- b Granulocytes $< 50\%$ of differential count (may be completely absent)
- 2 Anemia absent (cf aplastic anemia leukemia etc)

Treatment

A Emergency Measures

- 1 Discontinue all unnecessary medication and always discontinue offending drug promptly
- 2 Control of infection during the period of profound neutropenia
 - a Antibiotics May be used prophylactically to prevent infection or therapeutically to control infection when it develops. The chief disadvantage of routine prophylactic use of broad spectrum antibiotics (tetracyclines chloramphenicol etc) is that if infection develops the causative organism (especially the staphylococcus) may be resistant to a wide variety of antibiotics preparations. Therefore it is advisable under hospital conditions to withhold antibiotic therapy until infection develops then isolate the causative organism perform antibiotic sensitivity tests and administer the antibiotic to which the organism is most sensitive. During the period required to culture the organism and perform the sensitivity tests (48 hours or more after the first signs of infection have developed) a broad spectrum antibiotic should be given. For details concerning the selection and administration of antibiotics see page 514.
 - b Sulfonamides (auton) may be employed if appropriate antibiotic preparations are not available providing sulfonamides are not the offending drug.
- 3 Granulocyte formation. Purine nucleotide, folic acid, vitamin B₁₂ and pyridoxine are of doubtful value in producing leukopoiesis.

B General Measures

- 1 Hospitalize and isolate from avoidable infections
- 2 Bed rest
- 3 Careful skin and oral hygiene
 - a Saline mouth wash
 - b Local treatment of skin and mucous membrane ulcers (see page 96)
- 4 Nourishing liquid or soft diet as tolerated by painful buccal lesions. Parenteral feeding may be necessary (see page 25)
- 5 Follow patient blood status by serial blood counts

Prophylaxis

A patient receiving a drug capable of producing agranulocytosis must have regular and frequent WBC during the course of the therapy. The patient should also be cautioned to discontinue the drug promptly in the event of sore throat skin rash chills or fever and to report to his doctor at once.

LEUKEMIAS

Leukemia is an acute or chronic invariably fatal disease (of group of disease ?) of unknown etiology in living the hemopoietic tissues and usually giving rise to abnormal leukocytosis and associated anemia. Immature leukocytes are usually present in the peripheral blood although there may be no alteration in the blood count or there may be merely an increase in the number of normal leukocyte.

ACUTE LEUKEMIA (code No 50 7921)

Diagnosis (See table on page 234)

Treatment

Treatment is only palliative although this may change with the new chemotherapeutic agents. Radiation therapy is usually of more harm than benefit.

A Combat anemia and bleeding tendency by repeated blood transfusions as indicated. Improvement is usually transient although short remissions have been reported.

B Antimetabolite Therapy

1 Folic acid antagonists (Use with caution.)

a Agents most commonly used

(1) Aminopterin (4 aminopteroylglutamic acid)

(2) Methotrexate (formerly called amethopterin) (4 aminomethylpteroylglutamic acid)

b Indications for use. Acute leukemia especially in childhood. Some degree of improvement may be expected in approximately two thirds of the children treated and marked improvement will occur in about 50%. In children with acute leukemia the remission rate obtained with folic acid antagonists is much lower (12 to 15%).

c Dose. Folic acid antagonists may be given orally or intravenously in the form of intravenous infusion. The size of the dose should be gauged by the age, weight and physical condition of the patient. Remember that the range between the therapeutic and the toxic dose is very slight. Patients receiving these drugs must be watched closely and frequent blood count and bone marrow examinations made. Treatment must be discontinued promptly if toxic symptoms appear, if pancytopenia and hypoplasia of the bone marrow develop.

(1) Aminopterin Children 0.5 to 1.0 mg (1/20 to 1/50 gr) daily administered 1.0 to 2.0 mg (1/50 to 1/30 gr) daily.

(2) Methotrexate Children 2.5 to 5.0 mg (1/24 to 1/12 gr) daily administered 5 to 10 mg (1/12 to 1/6 gr) daily.

d Toxicity. The most important toxic manifestation is stomatitis, diarrhea, ulcerative lesions at any site in or throughout the gastrointestinal tract, gingivitis, stomatitis, hemorrhage (sometimes massive) and pancytopenia associated with severe hypoplasia or aplasia of the bone marrow.

e Methods of minimizing toxicity (not always effective).

(1) Prompt discontinuation of folic acid antagonistic therapy.

DIFFERENTIAL DIAGNOSIS OF THE LEUKEMIAS AND RELATED DISORDERS

Di case	Duration	Spleno megaly	Hepato megaly	Lymph nodes	WBC		Bone Marrow
					Total Count (usual rang)	Differential	
1 Chronic granulocytic leukemia (code No 502 792)	36 mos (8 mos to 16 yrs)	+++	++	±	20 000 500 000	Immature myeloid cells	Myeloid infiltration
2 Chronic lymphocytic leukemia (code No 503 792)	42 mos (8 mos to 8 yrs)	++	+	++	30 000 100 000	Immature lymphoid cells	Lymphocytic infiltration
3 Chronic monocytic leukemia (code No 506 792)		±	±		20 000 100 000 (2 000 500 000)	Immature monocytic cells	Monocytic infiltration
4 Acute leukemias (code No 50 7921)	8 wks (2 wks to 6 mos)	±	±	±	15 000 30 000 (2 000 100 000)	Blast cells (often un differentiated)	Leukemic infiltration Blast cells may be difficult to differentiate
5 Aleukemic myeloid (code No 502 7923)		+	±	±	4 000 (1 000 6 000)	Immature cells (few)	Leukemic infiltration (not always)
Agnogenic myeloid metaplasia of spleen (code No 520 958)	11 yrs	+	±	±	20 000 50 000	Immature myeloid cells Nucleated etc.	Normal aplastic or hyperplastic marrow

Not

The anemia associated with the leukemias is usually normocytic and may vary from mild to severe. The platelets are usually increased in chronic granulocytic leukemia but may be decreased in all other leukemias; the platelet count is usually decreased as described.

- (2) Parent r l administration of Leucovorin C l ium
N N D (sy nthetic Leuco o toc itrovorum factor) in
a ratio of 1 uc vorin to a agonist of 1 : 10 : 1
- (3) Blood t ansf sio Spaced at intervals to maintain
RBC at 2.5-3.0 million/cu mm
- (4) Ant b i tics If inf cti n dev lops (use appropriat anti
biotic aft r causative o rganism has been isolat d and
s nsitiv ty tests p r formed)

2 Purin a t go lists These ag nt only recently introduced
are et il in an inv stigativ st g of developm nt but in
gen al th y appe r t be less toxic than th f li acid
antagonists

Ag ts available for us

- (1) Me ptopurin N N D (Purin thol® 6 M P)
- (2) 6 Chlo opu ine
- (3) O D a oacetyl l e ine (azas in)

b I dicatio s f r use Ac te le k m sp ially in adult
An initial r mi sio rate of approximat ly 55% h been
epo ted i adult and 50% in childre f l l wing t e t
m t with 6 m r ptopuri e Th stat s f hlo opurine
d zaseri remain u determ d at p s t

c D sag and pro ed re

- (1) 6 Me captopu ine (6 M P) 2.5-4.0 mg /Kg body
weight/d y
- (2) 6 Chlo pu ine 10-20 mg /Kg body w ight/day
- (3) Az e l e 2.0 mg /Kg body weight/day

This compound admin st ed orally in a ingle
do e or in a d vided do twi e daily The ange b tw n
th the ap utic and the toxic d e is wider than it is with
th f li acid antagon st and h n pu in ant gonit
appear t be om what saf to Me apt pu i e is
the drug of hoi e for i t l the py t s dminist d
d ly r g dless f th developm t of p ono n d d gr
f leukopenia and n utrop n a pan ytop until bl t
(st m) lls pr lymphocyt p grs ul ytes o pro
mono yt s virtually dis ppe f om th b ne m w or
until t ic sympt m d elop (s below) T e tment then
is di ontin d and the pati ti watch d l ely If
mplet lni l and h matol gi rem s ens no
furthe t tment is giv n until relap occu s If no
mission an in mpl t r mission i obtained th pa
ti nt i placed on maint nance th py (usu lly les than
2.5 mg /Kg b dy weight/day) In a es in whi h e let
an t 6 m rcaptopurin appe to b de loping f
v bl p om tim s an be obt ined by givi g
a ase in in omb tion w th 6 M P or by adminis
t ng 6 hlo opu i

Th ful u f p e ant g nust d pend upo
lose obs v tio of th pati t and n rial blood and
bone ma w xamin t o s Th bo m r w rath
than th p ipher l blood is employ d as th p in ipal
the apenti guid During th tag of drug indu ed bon
marrow upp si p pe ly spac d blood t an fu tions
m t be giv n as pp rti m assure

d Toxic ty Toxic manif stations of mild d gr har
d by ano xia and naus f qu tly e obs ved

More severe toxic manifestations include vomiting, stomatitis, diarrhea, melena and rarely fever. Treatment should be discontinued only if the more severe toxic manifestations develop. Suppression of hemopoiesis (pancytopenia, hypoplastic bone marrow) is not indicated as a toxic manifestation since it is employed as a therapeutic guide. Irreversible suppression of hemopoiesis has not been observed in patients treated with 6-mercaptopurine. Toxic manifestations usually disappear within 3 to 4 days after termination of the disease.

Stoddard ACTH. The compound will induce clinical and hematological improvement in 50-65% of children with acute leukemia, but the remission rate in adult cutaneous leukemia is low (about 12%). Unfortunately patients in whom temporary remission has been induced become refractory to subsequent courses of this drug (see page 423).

CHRONIC LEUKEMIAS (code No 50 792)

Diagnosis (See table on page 234)

Treatment

Treatment is only palliative. Think in terms of controlling all phases of the disease for as long as possible, but do not place emphasis on the attempt at combating leukocytes. Some patients require treatment for periods of years.

A Radtke on Leukemia Enlargement and Infiltration. Presure symptoms are arising from enlarged glands and viscera may produce mechanical disturbances in the pericardium, involvement of vital structures, involvement of vital structures with leukemic infiltration may occasionally require surgical intervention although the epithelial neoplasms.

1. Indication

X-ray. The form of treatment may be utilized says semi-theapy (tuberculin, adjuvant) or surgical. In all the apy (treatment) to produce lymphadenoma, enlarged spleen, nodular leukemia infiltration (etc). It must be determined only by trained radiologist.

b Radtke on lymphomas (P32). The form of infiltration is important for it is to total body infiltration and has the advantage of not causing infection. In known cases, however, it is valuable in medical treatment of equipment handling suitable.

2 Chemotherapy. The use of chemotherapy agents having an anti-leukemic effect is an effective method of treatment. Hospitalization usually is not required. Since the number of agents that will effectively control the clinical picture of leukemia is relatively small, the use of chemotherapy is greater than those available for treatment of chronic lymphocytic leukemia and in the method of procedure for the two diseases is different. The treatment will be discussed separately for each. Chronic lymphocytic leukemia. In the elderly patient, chronic lymphocytic leukemia is relatively benign and frequently not treated at all. The indication for

treatment are markedly enlarged lymph nodes especially if causing persistent symptoms anemia due to bone marrow infiltration or extensive leukemic infiltration of viscera skin etc

(1) Triethylene Melamine N N D (TEM) dispensed in 1 mg (1/60 gr) and 5 mg (1/12 gr) oral tablets. This relatively new drug appears to be the agent of choice because of its pronounced destructive effect on the mature lymphocyte. Use with caution especially if the leukocyte count is below 50,000 cells per cu mm. If the leukocyte count is in excess of 50,000 cells per cu mm give 5 mg (1/12 gr) of TEM together with 2 Gm (30 gr) of sodium bicarbonate orally 1 hour before breakfast (Sodium bicarbonate prevents reaction of TEM with substances in the gastrointestinal tract and permits absorption of the entire dose). On the following day give 2.5 mg (1/25 gr) of TEM plus 1 Gm (15 gr) of sodium bicarbonate 1 hour before breakfast. Then wait 1 week and check blood counts. Repeat the administration of TEM at weekly intervals reducing the weekly dose to 5 mg (1/12 gr) when the leukocyte count falls below 50,000 cells per cu mm and to 2 to 3 mg as the leukocyte count approaches normal. When normal leukocyte values have been attained discontinue TEM therapy. Remission may last from 6 to 24 months. How ever during remission blood examinations should be made at intervals of 1 or 2 months.

If the initial leukocyte count is above normal but below 50,000 cells per cu mm give 5 mg (1/12 gr) of TEM orally together with sodium bicarbonate once each week until the desired result is obtained.

(2) Nitrogen mustard (HN) Remission is obtained with nitrogen mustard in chronic lymphocytic leukemia usually are shorter than those obtained with TEM and therefore TEM is preferred to nitrogen mustard therapy. For details of administration of nitrogen mustard see page 241.

b. Chronic granulocytic leukemia. In contrast to chronic lymphocytic leukemia chronic granulocytic leukemia always should be treated at the time the disease is first discovered. Agent available for treatment is listed in red reference.

(1) Triethylene Melamine, N N D (TEM). Quite effective in controlling chronic granulocytic leukemia for long periods of time. It has the advantage of giving remission lasting from 3 to 10 months but due to the large dose employed it has the disadvantage of causing nausea and vomiting (sometimes severe) in some patients for several hours after administration. However nausea and vomiting may be minimized by administering 25 to 75 mg (3/8 to 1/4 gr) of chlorpromazine (Thorazine®) before and at 3 hour intervals after the administration of TEM. It is important to remember that the dosage schedule for TEM in chronic

granulocyte leukemia is significantly higher and therefore quite different from that used in chronic lymphocytic leukemia.

If the leukocyte count is in excess of 50,000 cells per cu. mm. give 10 mg ($\frac{1}{8}$ gr.) of TEM and 2.4 Gm (30.60 gr.) of sodium bicarbonate orally 1 hour before breakfast. On the following day give 5 mg ($\frac{1}{12}$ gr.) of TEM and 2 Gm (30 gr.) of sodium bicarbonate 1 hour before breakfast. Repeat the above procedure at weekly intervals after first performing blood counts. Reduce weekly dose of TEM when the leukocyte count falls below 50,000 cells per cu. mm. and discontinue therapy when leukocyte values approach normal.

If the initial leukocyte count is below 50,000 cells per cu. mm. start with 10 mg ($\frac{1}{8}$ gr.) of TEM weekly. If response to this dose is unsatisfactory give 12.5 to 15 mg ($\frac{1}{5}$ to $\frac{1}{4}$ gr.) each week.

- (2) **Bulfan** Δ N D (Myleran[®]), dispensed in 2 mg ($\frac{1}{30}$ gr.) tablets for oral use. Utilize only in patients having chronic granulocytic leukemia with high leukocyte count. Do not use in patients with normal or subnormal leukocyte counts. Give 3.5 mg ($\frac{1}{15}$ to $\frac{1}{8}$ gr.) daily by mouth until maximum hematologic improvement achieved, performing blood counts every second third day. Thrombocytopenia of serious degree may develop on daily oral doses of 10 mg ($\frac{1}{6}$ gr.) or more. When the leukocyte count has returned to a normal level, place the patient on a maintenance dosage of 2.4 mg ($\frac{1}{30}$ to $\frac{1}{15}$ gr.) daily. Myleran is ineffective in acute leukemia and in chronic lymphocytic leukemia.

- (3) **Mercaptopurine** N & D (Purinethol[®]), 50 mg ($\frac{3}{4}$ gr.) tablets for oral use. Give 3.5 mg ($\frac{1}{20}$ to $\frac{1}{12}$ gr.) /Kg./day in a single or equally divided dose by mouth until the leukocyte count approaches normal. Maintain the therapy (2.5 mg /Kg./day or slightly less) than is required to control the disease. 6 M.P. is ineffective in chronic lymphocytic leukemia.

- (4) **Urethane**[®] (ethyl carbamate), dispensed in 0.5 Gm. ($\frac{1}{2}$ gr.) plain or enteric coated tablets for oral use. This compound will control chronic granulocytic leukemia for relatively long periods of time but it has the disadvantage of causing nausea and anorexia in a high proportion of patients and vomiting in some. Give 0.5 to 1.0 Gm. ($\frac{1}{2}$ to 1.5 gr.) t.i.d. until leukocyte count returns to normal, then place on maintenance therapy giving the smallest amount necessary to keep the leukocyte count at or near normal levels.

- (5) **Powder solution** (Potassium Asenite Solution, N.F.) may be of value when radiation therapy is contraindicated or unavailable.

For the choice of administration begins with an initial dose of 0.3 cc (5 gtt. or 5 min.) t.i.d. orally for 2 days. This dose is increased by 0.05 (1 gt. or 1 min.) every other day until a dose of 0.6 (10

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If the leukocyte count is in excess of 50,000 cells per cu mm, give 10 mg ($\frac{1}{8}$ gr) of TEM and 3.4 Gm (30.60 gr) of sodium bicarbonate orally 1 hour before breakfast. On the following day give 5 mg ($\frac{1}{12}$ gr) of TEM and 2 Gm (30 gr) of sodium bicarbonate 1 hour before breakfast. Repeat the above procedure at weekly intervals after first performing blood counts. Reduce weekly dose of TEM when the leukocyte count falls below 50,000 cells per cu mm, and discontinue the drug when leukocyte values approach normal.

If the initial leukocyte count is below 50,000 cells per cu mm, start with 10 mg ($\frac{1}{8}$ gr) of TEM weekly. If response to this dose is unsatisfactory, give 12.5-15 mg ($\frac{1}{5}$ to $\frac{1}{4}$ g) each week.

- (2) **Buulf** (N N D) (Myel-an[®]) dispensed in 2 mg ($\frac{1}{30}$ gr) tablets for oral use. Useful only in patients having chronic granulocytic leukemia with high leukocyte counts. Do not use in patients with normal or subnormal leukocyte counts. Give $\frac{1}{15}$ to $\frac{1}{8}$ gr ($\frac{1}{15}$ to $\frac{1}{8}$ gr) daily by mouth until maximum hematology improvement is achieved performing blood count every second or third day. Thrombocytopenia of serious degree may develop on daily doses of 10 mg ($\frac{1}{8}$ gr) or more. When the leukocyte count has returned to a normal level, place the patient on maintenance dose of 2.4 mg ($\frac{1}{30}$ to $\frac{1}{15}$ g) daily. Myel-an is ineffective in acute leukemia and in chronic lymphocytic leukemia.
- (3) **Mercaptopurine** (N N D) (Purinethal[®]) 50 mg ($\frac{3}{4}$ g) tablet for oral use. Give 3.5 mg ($\frac{1}{20}$ to $\frac{1}{12}$ gr) /Kg /day in a single or equally divided doses by mouth until the leukocyte count approaches normal. Maintenance therapy (2.5 mg /Kg /day or slightly less) than is required to control the disease. S.M.P. is ineffective in chronic lymphocytic leukemia.
- (4) **Urethane**[®] (thyl arbamat) dispensed in 0.5 Gm ($7\frac{1}{2}$ gr) plain orange colored tablets for oral use. This compound will control chronic granulocytic leukemia for initially long periods of time, but it has the disadvantage of causing nausea and anorexia in a high proportion of patients and vomiting in some. Give 0.5 to 1.0 Gm ($7\frac{1}{2}$ to 15 gr) tid until leukocyte count returns to normal, then place on maintenance therapy giving the smallest amount necessary to keep the leukocyte count near a normal level.
- (5) **Fowler's solution** (Fowler's Antacid Solution N.F.) may be of value when radiation therapy is contraindicated or unavailable.

Forker's technique of administration begins with an initial dose of 0.3 cc (5 gtt or 5 min) tid orally for 2 days. This dose is increased by 0.05 cc (1 gtt or 1 min) every other day until a dose of 0.8 cc (10

gtt or 10 min) t i d is reached Further incr a e of d e 0 05 cc (1 gt o 1 min) daily until toxic symptoms occur (anore : nausea and vomiting diar rhe) or the leukocyt count appoa hes normal Ds continue the drug for 2 5 days and then d crease the maximum dose by 0 05 cc (1 gt or 1 min) daily until a maintenanc l vel t 0 3 0 5 cc (5 gtt r 5 min t 8 gtt or 8 min) t i d is rea h d This do is con tinued indefinitely keep ng the patient unde c rel l observation

(6) Nitrogen mustard (HN₂) May p odu full clinical remissions in c tain early and m derately advanc d cases of chronic granulocytic l ukemia these are s m lar to x ray response but are of short r duration Nitrogen mustard is not recomm nded for a ute leuk mia (ee page 241)

B Treatment of Certain Hematologic Abnormaliti s

1 Anemia Dete mine whe the o not the an mia is myel ophthisic or hemolytic

a Myelophth s T eat w th the app opriat anti l uk mic chemotherapeutic agent Adequ t nutr tion including supplementary v tamins is important but the adminis tration of n alts is of no value P riodic t ansfusions of whole blood may be ne es ry until the d sir d hemo therapeut c sult has been attain d

b Hemolyt Tr at w th st ro ds or c rt cotropin (ACTH) (see p g 228) If th h molyt c anemia of chron c le kemia nn t be ontr l d by hormone therapy spl n tomy may be n s ry

2 Bleeding tendencies Pu puric and hemorrhagic ph nome a a e often due t th asso iated thrombocytopenia Trans f si ns of fresh whol blood are indicated Tolu dine blue rep ted to be of valu

3 Hem lyti crises S p g 228

C Oth Symptomatic M a u s

1 Treatm t of p ur i S e p ge 66

2 Treatm nt of ulce t i ve t matitis See p g 261

LYMPHOMAS (code No 820) and LYMPHOSARCOMAS (code No 821)

A large ll defined gr p f di ease har t rized by pro gr ss e p olif ration of th hematopoi t t ssu s and manif st d by var ble inv lvement f lymph n d s splee bon mar ow l ve and other reti loendotheli l stru tu tog the with c n st tut onal symptoms of feve w ight l ss h mo h gic tendencies and an mia Th ex ct lte lationship f th d sea s are not known ther f re all cl ssif cati s rem i arbitrary and cont o versial Clin al typ s re oft n ind finite and m y merg into one another

Tre tment

Ce tain gen ral principles of manag me t may apply to th se diseases as a group

A General Measures Measures directed toward maintaining optimum general health should be carried out both as a means of influencing the course of the disease and as a means of preventing complications.

B Radiation and Drug Therapy The effects of radiation and certain chemotherapeutic drugs may be palliative or curative but results are often equivocal. The susceptibility to a specific therapeutic agent and the duration of effectiveness (for both palliation and cure) will vary not only with the disease but also with the stage of the disease. Previous therapy and the response of the patient to patient. The table on page 238 outlines the response of the various histologic types to radiotherapy and chemotherapeutic agents.

Although clinical experience has shown that certain of these treatments are more amenable to therapy than others, final allocation must rest upon a trial of the therapy.

HODGKIN S DISEASE (code No 550 954)

A Pathogenesis A disease invariably fatal, characterized by a granulomatous (lymphomatous?) disease of unknown etiology involving the lymphoid tissues of the body. It is manifested by progressive enlargement of lymph nodes, spleen, and other lymphoid structures and constitutional symptoms of fever, weight loss, and anemia. The lesions can involve any and all tissues throughout the body. The disease is protean. Several clinical and pathological types are recognized, ranging from a usually more benign form, paraneoplastic, with a survival time of 3 or more years, to a rapidly fatal form, sarcomatous, with a survival time of less than 1 year. The diagnosis is confirmed by biopsy and confirmed by biopsy to differentiate the condition from the other infectious granulomas and from the other lymphomas.

Treatment

A Definitive Measures (No known specific therapy is available.)

- 1 Local Therapy At present, local or total body irradiation probably is the palliative measure of choice. Clinical improvement is attained but regression is often incomplete. The disease is usually fatal and in no way does it alter the course of the disease. The average survival time is probably unchanged but the patient is made more comfortable. Unfortunately, if the disease progresses very effectively after subsequent courses of x-ray therapy, nitrogen mustard therapy would be indicated in radiorefractory patients.
- 2 Combination and Intragastric Mustard Therapy may also be a valuable benefit but is limited by its method of administration.
- 3 Nitrogen Mustard [methyl bis(2-chloroethyl)amine hydrochloride] (HN_2) HN_2 is present in the nitrogen mustard must be carefully employed. The indication is for the use of a widely disseminated chronic granulomatous Hodgkin's disease that has been refractory to therapy.
- b Chronic Granulomatous Disease Hodgkin's disease with visceral involvement (especially lung parenchyma).

Hodgkin's disease failing to respond to therapy.

irradiation

HN_2 should be administered only in a hospital and only by a physician experienced in its use this is because of the complications that may arise as a result of severe toxic reactions. The response is similar to roentgen irradiation but the remission is shorter (usually 1 to 3 months). HN_2 is given I V in a single dose. The average dose is 0.4 mg (1/160 gr)/Kg body weight but slightly more or less than this amount may be desirable in certain cases (range 0.3 to 0.8 mg/Kg). HN_2 is stable in dry form but unstable in aqueous solution. Therefore fresh solutions must be used. Add 10 cc (2 1/2 dr) of sterile isotonic sodium chloride solution to 10 mg (1/8 gr) of the dry salt in a sterile container. Calculate amount the patient needs and draw this quantity into a sterile syringe of suitable size. Inject slowly taking no less than 10 minutes for the full amount into the stream of an infusion of saline or glucose solution that is being delivered at a rapid rate. Do not inject directly into a vein. Great care should be taken to prevent the HN_2 solution from coming into contact with the skin or escaping from the vein into the surrounding tissues.

- 4 Triethylene Melamine N N D (TEM) (See page 238) Mild cases of chronic granulomatous Hodgkin's disease sometimes can be controlled for months or longer by the oral administration of 50-75 mg (1/12-1/8 gr) of TEM (give with 2 Gm (30 gr) of sodium bicarbonate 1 hour before breakfast) once each month. However TEM is more valuable in treating lymphoma than Hodgkin's disease.

An important use of TEM is intrapleural administration in patients with serous or chylous pleural effusions. After withdrawing as much pleural fluid as possible 5 mg (1/22 gr) of sterile powdered TEM is dissolved in 5 cc (1 1/4 dr) of distilled saline solution and injected into the pleural cavity. After withdrawal of the thoracocentesis needle the patient is asked to change his position every 5 minutes for 30 minutes so that the TEM comes in contact with extensive areas of visceral and parietal pleura. This procedure is even more effective in treating serous or chylous pleural effusions in lymphoma than a smaller dose (2.5 mg) of TEM should be given.

- 5 Surgical excision. Wide surgical excision may be indicated for localized lesions especially if it is primary (isolated or presenting area). This is a debatable measure.

B General Measures

- 1 Maintaining good living hygiene with adequate diet, exercise and rest.
- 2 Hospitalization is recommended during febrile phases with other complications of the illness.
- 3 Transfusions of whole blood and other supportive measures should be instituted as the various medical indications arise.

MULTIPLE MYELOMA (code No 533 8221)

A chronic disease of unknown etiology usually occurring after 40 years of age marked by circumscribed or diffuse proliferation

hyperplasia of plasma cell and characterized by neuralgic and skeletal pains spontaneous fractures x-ray evidence of skeletal destruction Bone Joints proteinuria anemia renal insufficiency and an invariably fatal termination

Treatment

A. Definitive Measures

1. X-ray therapy may provide symptomatic relief but it is doubtful if it significantly alters the course of the disease
2. Urthanes® (thylca bromide) Urthane® in doses of 1.5 to 6.0 Gm (22½ to 90 gr) per day as tolerated may provide symptomatic relief but probably does not prolong life. Careful follow-up with periodic leukocyte counts is necessary
3. Stilbamidine diethiodate N N D Formally used to relieve severe skeletal pain in relation to x-ray therapy but since it causes severe intracerebral edema and neurological gain a significant proportion of cases it now is used infrequently in the treatment of myeloma. When used it is given in freshly prepared solution containing 150 mg (2½ gr) stilbamidine I.M. or I.V. daily or very other day for from 8 to 50 injections. The patient should be placed upon a diet low in animal protein. Careful assessment of renal function prior to and during drug therapy is necessary. Although patients may have temporary clinical improvement the course of the disease remains progressive and fatal

B. General Measures

1. Permit optimum general health by adequate diet and sleep
2. Caution patient against exposure to undue physical trauma because of susceptibility to fracture
3. Whole blood transfusions needed if anemia
4. Angiostatic pain
5. Encourage fluid intake in good urine output

BLEEDING DISEASES

The mechanisms involved in the maintenance of the coagulability and fluidity of the blood are as yet incompletely understood. The field is still a point concerning the blood coagulation mechanisms have failed to explain many phases of bleeding phenomena. The three bleeding mechanisms are (1) clotting defect (2) thrombocyte (platelet) defect and (3) capillary defect. They are all closely interrelated and poorly understood in many bleeding diseases. From a clinical standpoint it is perhaps more important to determine the total globulin for the bleeding. It is important to remember that the various individual hemostatic test may be limited to a wide variety of disorders and are seldom pathognomonic.

For simplicity of presentation and for purposes of familiarity however the following table of bleeding diseases is based upon the old classification. One should bear in mind that each diagnostic group is a truly difficult to establish in terms of absolute definition of specific coagulation factors.

BLEEDING DISEASES SUMMARY OF DIAGNOSIS AND TREATMENT

Diagnosis	Clinical Features	Diagnosis	Treatment	Prognosis
Congenital Hemophilia	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor VIII deficiency	Factor VIII concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor IX deficiency	Factor IX concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor X deficiency	Factor X concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor XI deficiency	Factor XI concentrate	Good
Acquired Hemophilia	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor VIII deficiency	Factor VIII concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor IX deficiency	Factor IX concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor X deficiency	Factor X concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Factor XI deficiency	Factor XI concentrate	Good
Vitamin K deficiency	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Vitamin K deficiency	Vitamin K concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Vitamin K deficiency	Vitamin K concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Vitamin K deficiency	Vitamin K concentrate	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Vitamin K deficiency	Vitamin K concentrate	Good
Liver disease	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Liver disease	Liver transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Liver disease	Liver transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Liver disease	Liver transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Liver disease	Liver transplant	Good
Kidney disease	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Kidney disease	Kidney transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Kidney disease	Kidney transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Kidney disease	Kidney transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Kidney disease	Kidney transplant	Good
Bone marrow failure	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Bone marrow failure	Bone marrow transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Bone marrow failure	Bone marrow transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Bone marrow failure	Bone marrow transplant	Good
	Long bleeding time Normal clotting time Normal bleeding time Normal clotting time	Bone marrow failure	Bone marrow transplant	Good

Treatment.A Prevent From Sources that Aggravate Bleeding (Tocantins)

- 1 Limit activities. Advise occupations, sports, or other activities which involve minimal physical hazards.
- 2 Prevent areas of body which are subject to injury
 - a Lubricate nostrils and other superficial bleeding sites with petroleum to prevent drying and cracking of scabs.
 - b Apply protective bandages, splint or casts to existing wounds to prevent repeated hemorrhages.
 - c Bandage lower extremities carefully to supply support to surface skin vessels when indicated.
- 3 Surgical procedures. Limit the number and extent of operative procedures to a minimum.
 - a Considered for elective surgery carefully.
 - b Minimize trauma, extent, and duration of operative procedures.
 - c Perform operative procedures in stages (e.g., extract one tooth or remove one tonsil at a time).
 - d Prepare patient for surgical procedure by appropriate hemostatic techniques (e.g., preoperative fresh whole blood transfusion or vitamin K).
- 4 Correct intrinsic factors
 - a Treat cardiac failure, hypertension when present.
 - b Correct symptoms of violent coughing, sneezing, etc.

B Local Bleeding Must Be Treated Promptly

- 1 Bandaging. Do not use dressings for hemorrhostasis.
 - 2 Topical thrombin may be applied locally for hemostasis.
 - 3 Thromboplastin. Less effective than thrombin.
 - 4 Adhesive. Adhesive Bandage, U.S.P. (Gelfoam® Oxyel® Fibrinogen®).
 - 5 Electrocoagulation.
 - 6 Chemical astringents. Use only of value only for small bleeding sites. Gipsix (see page 111). Use triethylrattadefichloride, tannic acid, or hemobad.
 - 7 Snake venom (Rusell Viper) 1:10,000.
- C General Hemorrhage or Generalized Bleeding. Must be treated by measures which combat shock, coagulable bleeding and correct anemia.
- 1 Combat shock (see page 27). Fresh whole blood (not older than 3-4 hours) is preferred because of its hemostatic as well as anti-hemodynamic effect. Plasma may be used when whole blood not available.
 - 2 Control bleeding
 - a Blood clotting tests
 - (1) Fresh whole blood transfusion. I.V. (see page 27 under Shock). The plasmalet plasma in refrigerated whole blood clot so at largely within 12 hours at the highest plasma hemofiltrate may permit muhlagpeiod. Blood transfusion effectively administers should be considered as an emergency support in all forms of hemorrhage regardless of cause.
 - (2) Plasma transfusion. Transfused off fresh frozen plasma (not older than 10 days) provides prothrombin

fibrinogen and hemophilic globulin and certain other factors which may be of value in controlling bleeding. There are no platelets in plasma.

- (3) Antihemophilic Globulin U S P (Cohn Fraction I) in average doses of 200 mg (sometimes up to 600 mg may be required) in 5-10 cc physiological saline causes a decrease in the spontaneous clotting time of the blood of hemophilic patients.

b. Vitamins

- (1) Ascorbic Acid U S P See page 63
 (2) Phytonadione U S P (vit. K₁ emuls. on M. phyto.)
 For use in acute hemorrhagic emergency due to hypoprothrombinemia. Give 50-100 mg I V very slowly.
 (3) Vitamin P and related compounds. Experimentally this group of agents has been reported to increase the capillary resistance in certain diseases in humans but clinical studies have been discouraging. Two preparations are Rutin N F 20-50 mg q.i.d. and asperidine methyl chalc. 50 mg q.i.d.

- c. Corticotropin (ACTH) and the steroids (see page 423) may produce striking remissions of the purpuric hemorrhagic reaction (increased red cell count and platelets and decreased bleeding tendency) and at least tide the patient over until a chance time as other more specific measures (e.g. transfusions, surgery) can be safely instituted.

- d. Antiheparin agents. In anaphylactoid shock secondary thrombopenic purpura, irradiation reaction, nitrates, mustard therapy, leukemia, menorrhagia and in certain other conditions heparin or a heparin-like substance is liberated in excess and appears to be responsible for a bleeding tendency (hyperheparinemia). The excess of heparin may at times be counteracted by the use of two agents, protamine sulfate or toluidine blue, which inactivate heparin by forming a bi-chemical complex.

- (1) Protamine Sulfate Injection N F 50 mg in 5 cc aqueous solution I M every 4-6 hours until platelets cease to appear. 150 mg in 250-500 cc of 5% glucose or normal saline given slowly I V is often given at the time of the final I M injection. Protamine treatment (see J A M A 139:1251, 1949) may provide a diagnosis as to the cause of the bleeding.

- (2) Toluidine blue 6-8 mg /Kg body weight dissolved in normal saline given slowly I V (over a 2 hour period) daily for 3 days and followed by 2-4 mg /Kg for 3 additional days. In preparation of the dye solution it must be passed through a Searle filter for sterilization and removal of large dye particles. Transient nausea and vomiting, bluish tint to skin and blue coloration of urine and feces may be encountered.

- e. Splenectomy. Removal of the spleen may be indicated in selected cases of primary thrombocytopenic purpura and in cases of secondary thrombocytopenia due to splenic diseases (hypersplenism). Demonstration of megakaryocytic activity by the bone marrow is essential to the proper evaluation of the individual. In general

spl enectomy is advised only in the "hypersplenic" forms of th ombocytopenic purpura (primary or secondary) i e. Guch, Banti's and granulomatous diseases of the spleen) but the operation may be indicated in very selected cases of bone marrow megakaryocyte deficiency. The decision as to need for splenectomy should be made by a trained hematologist.

Rh FACTOR

(Reaction due to Blood Transfusion code No. 010 38x)

When blood containing the Rh factor (from Rh positive donor) is introduced into a person without the factor (Rh negative) the Rh factor acts as an antigen and agglutinins may develop against it (anti Rh agglutinins). After the agglutinins have developed Rh-positive blood from donors is no longer suitable for transfusion. The agglutination and hemolysis of the donor cells is likely to occur. The severity of such transfusion reaction (as with intergroup reactions) may vary considerably. Rh sensitization is well known to develop by multiple pregnancies in Rh-negative women with Rh-positive husbands.

Precautions

A General Rules

- 1 All blood for transfusions should be Rh typed in addition to conventional intergroup typing and then cross matched with recipient blood.
- 2 Rh-positive individuals may safely receive blood only from Rh-positive donors.
- 3 Rh-negative individuals may safely receive blood only from Rh-negative donors.

B Specific Rules Never give Rh-positive blood to any of the following:

- 1 Rh-negative individuals who have had previous transfusions
- 2 Rh-negative women who have had multiple pregnancies by Rh-positive husbands
- 3 Rh-negative pregnant women
- 4 Infants with erythroblastosis

Testing

See under transfusion reactions on page 249

BLOOD TRANSFUSION

Physiological Reaction:

Blood is given in order that

- 1 In case of circulating fluid volume
- 2 In case of oxygen carrying capacity of blood
- 3 In case of protein concentration
- 4 In case of coagulability of blood
- 5 In case of immune bodies

Contraindications

Transfusions must be given carefully in cases of acute pulmonary edema, cardiac failure, nephritis, and pulmonary embolism or infarction.

Preparation for Blood TransfusionsA. Typing and Cross matching

1. Determine blood type of recipient. Use known typing sera: Anti A Blood Grouping Serum U S P (or serum from Type B blood) and Anti B Blood Grouping Serum U S P (or serum from Type A blood). Blood type may be determined according to chart below.

Recipient's + b c		Recipient's Type	
Anti B Serum	Anti A Serum	Landsteiner	Moss
+ Agglutination	+ Agglutination	AB	I
0 Agglutination	+ Agglutination	A	II
+ Agglutination	0 Agglutination	B	III
0 Agglutination	0 Agglutination	O	IV

2. Donor should always be of the same blood type as recipient. Cross match as indicated below. In emergency situations, Type O (Moss IV) blood (universal donor) may be administered to any type. Type AB individuals may receive blood of any type (universal recipient).
3. Always perform direct compatibility test between donor and recipient blood before each transfusion, even if the blood came from a previously compatible donor. This is done by mixing recipient's cells (RC) and donor's serum (DS) on one side of a glass slide, and donor's cells (DC) and recipient's serum (RS) on the other side.

Donor's cells
+
Recipient's serum



Recipient's cells
+
Donor's serum

The slide is rocked back and forth for 5 minutes and examined with the low power microscope. If there is any agglutination or suggestion of hemolysis, a new donor must be found.

4. Whenever possible, determine the Rh of the recipient. Rh negative recipients should receive only Rh negative blood. Rh positive recipients may receive Rh positive or Rh negative blood in emergency when no compatible Rh positive blood is available.

B. Disease Which May Be Transmitted By Blood Transfusion

1. Syphilis. Donor should always have a serological test for syphilis.
2. Malaria and hepatitis. Blood from a person with a history of malaria or infectious or homologous serum hepatitis should not be used.

Technic of Blood Transfusion

There are two methods for administration of blood (1) indirect transfusion using modified blood (blood to which anticoagulants have been added) and (2) direct transfusion (blood transfused directly by vein without addition of any substance). The first method is now used almost exclusively.

A Indirect Transfusion Using Modified Blood Citrate is used most frequently as the anticoagulant.

1. Collection of blood. A specially prepared vacuum flask that contains sodium citrate-citric acid in dextrose solution is commonly used to collect 500 cc (1 pt.) of blood. The

collection apparatus is equipped with a valve that allows the amount of suction to be regulated. This is the technique used in most blood banks.

2. Administration. The collecting bottle is connected with a Y tube to a small bottle of saline. The blood bottle is clamped off at one arm of the Y. The saline is then used to fill the tubing and start the infusion. After the saline has begun to flow into the vein, the blood bottle is allowed to flow and the saline stopcock at one of the arms of the Y tube.

B Direct Transfusion This technique uses an apparatus consisting of a large syringe and a smooth working 3-way stopcock. The blood is drawn into the syringe from the donor through the stopcock, turned, and the blood injected immediately into recipient.

Precautions in Administration

- A. Always administer alkali (5 Gm. or 75 gr. sodium bicarbonate) orally. 250 cc ($\frac{1}{2}$ pt.) M/6 sodium lactate buffer beginning transfusion. Prophylaxis for hemolytic reaction.
- B. Never warm blood before administration.
- C. A rate of 40 to 50 drops per minute or 150 cc (5 oz.) per hour. Can be given at maximum rate of 1 (15%) per second.
- D. In cases with myocardial insufficiency give about 1 cc (15%) per minute (12 to 15 drops per minute). Never give over 75 (2½ oz.) in 1 hour except in the most critical cases.

COMPLICATIONS OF TRANSFUSION

Transfusion Reaction

Transfusion should be stopped immediately if patient complains of chills, general itching, sensation of anxiety, pruritus, oppression, pain in back of neck, thorax and lumbar area, or a feeling of fullness of the head.

- A. Hemolytic Reaction. Most severe of all and may be fatal. Symptoms in tension develop usually appear during the transfusion immediately afterward. Hemolytic reactions almost always caused by incompatibility of blood.

B. Allergic Reaction. Usually occurs following transfusion.

1. Mild form. Urticaria, general edema and sinusophiles.

2. Moderate form. Difficulty of breathing, a feeling of constriction, itching, pruritus, and pharyngeal edema.

- C. Chills and Rigors (Pyrexia). Most common reaction. Reaction usually occurs within 15 minutes to 1 hour after transfusion. Characterized by chills, rigors, followed by fever.

Treatment of Transfusion Reactions

A Hemolytic Reactions

- 1 Rationale To attempt to prevent the precipitation of acid hematin in the renal tubules. Therefore alkalinization of urine and forcing of fluids is important
- 2 Definitive measures
 - a Give 10 Gm (150 gr) sodium bicarbonate orally at once and every 4 hours. If patient is unable to void 10 to 20 Gm (150-300 gr) sodium bicarbonate (specially prepared see page 15) in 100 cc of distilled water I.V. or 500-1000 cc (1-2 pt) of M/6 sodium lactate I.V. Repeat the dose in 8 hours or sooner if the urine becomes acid
 - b Collect all urines and examine for hemoglobin. Continue alkalinization until no further hemoglobin is present
 - c Supply fluids orally or by parenteral means to maintain urine volume of at least 1500 cc (1 1/2 qt) per 24 hours as long as renal function is normal (See acute renal failure page 303)
 - d In severe or repeated hemolytic reactions where repeated transfusions may be necessary corticotropin (ACTH) or one of the steroids is indicated (see page 423)

B Allergic Reactions Treat as an allergic reaction

- 1 Give 0.2-0.5 cc (3-8 m) of epinephrine (adrenaline) 1:1000 subcut at once
- 2 If symptoms persist may try antihistamines (see page 45)

C Chemical Reactions

- 1 During chill keep patient warm by adding blankets and hot water bottles. This is usually all that is required
- 2 However since the differential diagnosis from the allergic reaction is often impossible give epinephrine (adrenaline) 1:1000 0.2-0.5 cc (3-8 m) subcut as soon as possible

Chapter 10

DISEASES OF THE GASTROINTESTINAL SYSTEM

NONSPECIFIC GASTROINTESTINAL SYMPTOMS

HALITOSIS (Bad Breath) (code No 619)

Halitosis can result from many causes and treatment is directed at removal or correction of these

Treatment

- A C orrect ba malit es of oral hygiene if patient
- B T e t E sting D s
 - 1 Chronic nasitis and sinus disease
 - 2 Dental caries gum infections tonsillitis infection etc
 - 3 Systemic diseases fevers and toxemias
 - 4 Chronic pulmonary disease e.g. lung abscess
 - 5 Gastrointestinal disease at any level of the GI tract
 - 6 Neuropsychiatric disorders where only the subjective complaint of "bad breath" presents
- C E l m i n te O ff nd g F o d s F r o m t h D i
 - 1 Q u l i c a n d s
 - 2 Rich or gassy foods if they are the known cause

HEARTBURN (Pyrosis) (code No 784.3)

Rule type if usual Consider specially disease of lower esophagus stomach biliary tract

Treatment

- A D r u g s
 - 1 A n t a c i d s These drugs (page 264) are often effective in relieving stomach although it is not felt that the relief is oblong and necessary rely depend upon the neutralization of the gastric hydrochloric acid
 - 2 S d a t a r e Antipruritic medication (see page 265)
- B B i d D i (See page 54)

NAUSEA AND VOMITING (Nausea code No 611) (Vomiting code No 614)

These symptoms may usually occur acutely and may be

due to a wide variety of psychic, reflex or central causes

- A Psychic Causes These are variable and may have either superficial or deep seated basis
- B Reflex Causes Disturbances of various gastrointestinal structures and other viscera are capable of exciting the vomiting center. Correction of this type of vomiting is therefore dependent upon removal or alteration of these reflex disturbances
 - 1 Irritation inflammation or mechanical disturbances at any level of gastrointestinal tract (from pharynx to rectum)
 - 2 Irritating impulses arising in any diseased viscera e.g. chol. cystitis
 - 3 Disturbances of semi circular canals e.g. seasickness
 - 4 Toxic action of cardiac drugs e.g. digitalis
- C Central (Vomiting Center) Causes
 - 1 Central emetics Emetine apomorphine morphine
 - 2 Exogenous and endogenous toxins
 - 3 Increased intracranial pressure
 - 4 Cerebral hypoxia Cerebral anemia or hemorrhage

Treatment

- A Acute Simple vomiting such as occurs following dietary indiscretion or as experienced in the morning sickness of early pregnancy may require little or no treatment. When necessary treatment consists of prescribing simple tolerated foods and occasionally mild sedative and antispasmodic drugs
- B Prolonged Severe or prolonged nausea and vomiting requires careful medical management. Specific causes must be corrected or eliminated. The following general measures may be utilized as adjuncts to specific medical or surgical measures
 - 1 Fluid and nutrition Maintain hydration and nutrition. Withhold foods by mouth temporarily. Administer 5-10% glucose in saline or water I.V. in quantities sufficient to maintain adequate hydration. When oral feedings are resumed commence with dry foods in small quantities e.g. salted crackers Graham crackers etc. With morning sickness these foods may best be taken before arising. Later change to frequent small servings of simple palatable foods. Hot beverages and clear broths and cold beverages iced tea and carbonated liquids (especially ginger ale) are tolerated quite easily. Avoid lukewarm beverages. Always consider patient's food preferences
 - 2 Drugs
 - a Sedative antispasmodic drugs may be helpful (see page 266)
 - b Ethyl aminobenzoate (benzocaine) 0.2 Gm (3 gr) with phenobarbital 20 mg ($\frac{1}{2}$ gr) every 6 hours p.r.n.
 - c Chlorpromazine Hydrochloride U.S.P. (Thorazine®) has been used for control of nausea and vomiting due to a wide variety of causes. It is administered orally in doses of 25-50 mg every 4 to 6 hours p.r.n. or orally in doses of 10-50 mg every 4 to 6 hours p.r. (see p. 40)
 - d Prochlorperazine N.N.D. (Compazine®) 5 mg i.i.d. q.i.d. orally when feasible or 5-10 mg (1-2 cc) deep into buttocks every 3-6 hours (not exceeding 40 mg/24 hours) has been reported to be valuable

e Meclizine Hydrochloride N N D (B namine®) 25 mg
daily may be of value in moderate cases

3 Psychotherapy

- Isolation of patient is recommended. Hospitalization may be necessary. Visiting should be restricted.
- Avoid unpleasant psychiatric stimuli such as strange odors, foul smelling or foul tasting medication, emesis basins or other unattractive objects as well as foods which are improperly prepared or served.
- Place patient on a definite treatment program and let it be known that something is being done. Hard-boiled or brutal techniques are to be avoided.
- Attempt to determine the psychodynamics of the nervous system and vomiting but avoid aggressive psychotherapy during acute phases of the illness.

HICCUP (Singultus) (code No 671)

Hiccup, although a common and usually benign symptom, may be a manifestation of any one of many diseases. It is important to rule out a wide variety of specific causes. Causes CNS disorders, pulmonary disorders, gastrointestinal disorders, renal failure, infectious diseases and other diseases.

Treatment

Treatment of the specific cause may suffice to relieve hiccup. However, it is usually necessary to use certain specific measures to provide relief from this symptom. Countless have been suggested for breaking the rhythmic reflex. All the treatment measures may fail and the symptom may become prolonged and severe as to jeopardize the patient's life.

A Simple Home Remedy These measures probably assist by diverting the patient's attention from the irritating stimulus, fight painful unpleasant stimuli or frightening patient from progressively paralyzing the pharynx (holding breath, pinching the nose, inhaling long fumigation, etc.).

B Drug and Medication

1 Sedation Any fifth valium, diazepam, USP, Pentobarbital, Sodium Barbital, USP, 0.1 Gm (1½ gr) orally 0.13 Gm (2 gr) by rectal proctitis.

2 Anesthetics Local anesthetics, e.g., cocaine may be applied to the pharynx. Gaseous anesthetics, e.g., nitrous oxide.

3 Antispasmodics Atropine, Sclaf, USP, BP 0.306 mg (1/100 1/100 g) may be given subcutaneously.

4 Amyl nitrite inhalation may be effective.

5 Carbon dioxide Hyperventilation with a plastic bag for 3 to 5 minutes, administration of 10-15% CO₂ mixture by face mask for 3 to 5 minutes.

6 Chlorpromazine or Hydrochloride, USP (Thorazine®) has been fully used for prolonged hiccup (see page 252).

• Surgical Measures Various pharyngeal operations, including esophageal atresia, pyloromyotomy may be indicated in certain cases.

cases which fail to respond to all other measures and which are considered to be a threat to life

CONSTIPATION (code No 630)

Eliminate specific causes of constipation first. Rule out colonic or rectal lesions, hypometabolism or psychogenic causes. Be especially suspicious of specific causes when there are sudden unexplained changes in bowel habits. Inadequate fluids and low residue diets may have a constipating effect. The following commonly used drugs which the patient may be receiving for an unrelated illness may cause constipation: belladonna and derivatives, narcotic drugs, bismuth salts, calcium salts, aluminum hydroxide gels (Amphojel®), aluminum phosphate gels (Phosphalge®) and iron salts.

Treatment

A. Correct Patient Attitude Toward Elimination

1. A daily bowel movement is not essential to normal health or well being. There is normally considerable individual variation in the frequency of bowel movements.
2. So called auto intoxication theories are unfounded.
3. Constipation particularly for short periods is seldom a cause for alarm.
4. Many symptoms (e.g., lack of pep) attributed to constipation have no such relationship.
5. Periodic purgation serves no tonic purpose.

B. Re-establishment of Regular Evacuation

1. The gastrocolic reflex should be utilized to optimal advantage by having patient set aside a regular daily period after a meal (preferably breakfast) for a bowel movement even when the urge to defecate is not present. This is based physiologically on the primitive reflex wherein distention of the stomach by food sets off a reflex evacuation of the colon. Explanation of the reflex vacuum as it occurs in infants after feedings appeals to many patients. Emphasize the fact that the normal reflex is preserved or lost by personal habits or social customs.
2. Sufficient time must be allotted to permit a leisurely performance of the act.
 - a. Patient may alter his daily schedule to permit more time for bowel movements.
 - b. Relaxation may be aided by reading a book, etc. while sitting on the toilet.
3. Cathartics and enemata should never be employed without direct advice or supervision of a physician if patient ever expects seriously to correct his constipation since these measures interfere with the normal bowel reflexes. For psychological reasons if not physiological it is sometimes inadvisable to discontinue such measures suddenly if patient has employed them for a prolonged period of time. It may be better to compromise temporarily with intermittent measures of bland laxatives and mild enemata (see next page). Chronic cathartic and enema addicts often defy all medical measures and their treatment is especially hopeless when

there are serious underlying psychiatric disturbances

C Diet In general the diet may be profitably modified to satisfy the following requirements (see pag 54)

- 1 Adequate volume Often constipation is merely due to inadequate food intake
- 2 Adequate bulk of residue This does not necessarily imply roughage such as bran Smooth or bland foods may be preferred in spastic constipation
- 3 Vegetable irritants Unless there is specific contraindication (e.g. intolerance) of sweet or raw fruits or vegetables may be of value in many cases of chronic constipation especially the so called atonic type
- 4 Adequate fluids The patient should be encouraged to drink adequate or large quantities of fluids other than reassembled water is available in the intestinal tract for passage of intestinal contents
 - a Six to 8 glasses of fluid per day in addition to fluid content of food are ordinarily sufficient
 - b The time honored glass of hot water taken a half hour before breakfast seems to exert a mild laxative effect

D Exercise Moderate physical exercise adjusted to individual needs and capabilities is essential Bed patient may profit by active and passive exercises Good tone of the external abdominal muscles is important Corrective physiotherapy may be employed in patients with ptotic abdomens

E Laxatives

- 1 Bland laxatives These agents should be employed temporarily during the bowel training (re-education) program or as a compromise measure in long standing cathartic and enemata addicts They are never intended as a substitute for a careful bowel training program They should be withdrawn as soon as the constipation improves

a Liquid Paraffin U.S.P. Liquid Paraffin B.P. (mineral oil) 15-30 cc (1/2-1) 1-2 times daily per os

b Agar U.S.P. B.P. with mineral oil 15-30 cc (1/2-1 oz) 1-2 times daily per os

Do not use mineral oil over prolonged periods in the intestine with absorption of foodstuffs particularly fat soluble vitamin There is also some risk of lipid pneumonia even from its oral use

Olive Oil, U.S.P. B.P. 15-30 cc (1/2-1 oz) 1-2 times daily per os

d Vegetable mucilage Pyllium Hydrophil Mucilloid N.N.D. (Metamucil[®]) 1-3 tsp b.d. to t.q. per os in full glass of water
Cassia Seed Gum Aromatic Fluid Extract U.S.P. 4 g c (1-2 d) h.s. very light

f Magnesia Magma U.S.P. (milk of magnesia) 15-30 (1/2-1) h.v. at night

g Sodium Phosphate U.S.P. B.P. (disodium phosphate) 4-8 Gm (1-2 d) in hot water before breakfast

h Docusyl Sodium Salt Inert U.S.P. (Cela[®] D₁ n₁) a u.s. weighting agent in combination with other laxatives
Young's 50 to 480 mg per day

F Enemas Enemas interfere with restoration of normal bowel reflex. These measures (as with above medications) should be utilized only as temporary means in chronic constipation or fecal impaction. They may be necessary for cleansing a bowel preparatory to diagnostic studies or proctocolysis.

- 1 Saline enema (non irritating) Warm physiological saline solution 500 2000 cc (1 pint to 2 quarts) p r n
- 2 Warm tap water (irritating) 500 1000 cc (1 pint to 1 quart) p r n
- 3 Soapsuds (S S) enema (irritating) 150 cc (5 oz) of soap solution n n 1860 cc (62 oz) water
- 4 Oil retent enema 180 cc (6 oz) of mineral oil or vegetable oil instilled in rectum in the evening and retained overnight. A cleansing soapsuds enema is given the following morning.

FECAL IMPACTION (code No 660 616)

This condition should be suspected in all severely constipated patients especially bed patients. Appropriate anticonstipation measures (see p 254) will usually prevent impaction.

Treatment

- A Manual removal of fecal impaction
- B Oil retention enema followed by cleansing enemas p r n (see above). Manual removal of impaction may be facilitated by this procedure.

FLATULENCE

Eliminate apathetic aus of flatulence Gastrointestinal gas is in large part due to swallowed air (aerophagia). However flatulence may be due to dietary causes and fermentation and organic disease of the digestive system.

Treatment

- A Correction of Aerophagia Anxiety states are often associated with deep breathing and sighing and the consequent swallowing of considerable quantities of air. When possible treat underlying anxiety features. A convenient and simple method for relieving the distention flatulence and belching during acute anxiety states with severe aerophagia is to have the patient place a small cork between his teeth while the effect of preventing air swallowing.
- B Correction of Physical Defects These sometimes interfere with normal swallowing and/or breathing.
 - 1 Structural deformities of the nose and nasopharynx e.g. nasal obstruction and adenoids
 - 2 Structural deformities of the teeth. Spitting defect
- C Good Hygiene and Eating Habit Instruct the patient to

- 1 Avoid eating too rapidly and too much
- 2 Avoid eating while under emotional strain
- 3 Avoid taking laxatives
- 4 Avoid chewing gum
- 5 Avoid dietary indiscretions

Diet

- 1 The diet should be composed of bland high protein low fat low carbohydrate foods (see page 54)
- 2 *Restrict gas-producing or irritating foods -*
 - a Avoid most raw fruits and vegetables especially cabbage cucumbers onion peppers celery tomatoes and beans
 - b Avoid sugar in large quantities or concentrated forms
 - c Avoid fried food
 - d Avoid nuts raisins berries and other seed fruits
 - e Avoid pices
 - f Avoid alcohol and alcoholic beverages

Medication These agents are of general unsatisfactory and attain a only of placebo value

- 1 Antispasmodic drugs (see page 266) are perhaps the most useful of the medications. Besides their antispasmodic effect they serve to diminish the flow of saliva (which is often excessive in the patients) thereby reducing the anorexia which accompanies the swallowing of the excessive quantity of saliva.
- 2 Solutio Peppermint U.S.P. B.P. 0.5 (1½ min.) t.i.d. in a small glass of water p.c.
- 3 Neostigmine Bromide U.S.P. B.P. (Pestigmin Bromide®) 15 mg (¼ gr) t.i.d. p.
- 4 Dehydrocholic Acid U.S.P. (Deholin®) 0.25-0.50 Gm (¾-3/4 gr) t.i.d. p.c.
- 5 Ox bile Extract N.F. (bile salt) 0.3 Gm (5 gr) t.i.d. p.
- 6 Adsorbent Kaolin N.F. B.P. and charcoal emulsion without rational physiological justification when one considers the quantity of gas to be absorbed as compared with the quantity of the drug to be given.

DIARRHEA (code No. 635)**Etiology**

Attempt to determine the etiology of an individual case when possible. The usual causes of diarrhea may be classified as follows:

- 1 Psychogenic diarrhea Nervous diarrhea
- 2 Gastrointestinal Achlorhydria
- 3 Intestinal
 - a Infectious a. Villenteritis amebiasis
 - b Enterocolitis Hemolytic enterocolitis
 - c Drug Catharsis habitus
 - d Stricture Gastroenteritis
 - e Neoplasms of the Colon
 - f Ischemic Colitis ulcerative colitis
- 4 Nutritional deficiency Spina
- 5 Primary diabetes Pancreatic insufficiency
- 6 Secondary diabetes Cholelithiasis duodenitis

- 7 Reflex from other viscera Pelvic pathology (extrinsic to GI tract)
- 8 Neurologic disease Tabes dorsalis
- 9 Metabolic disease Hyperthyroidism

Treatment

- A Eliminate the specific cause, whenever possible
- B Control Physiologic Change Induced by Diarrhea In addition to necessity for control of intestinal hyperperistalsis it is essential that the following secondary or complicating features be treated
 - 1 Fluid imbalance (dehydration) (see page 7)
 - 2 Mineral imbalance e.g. hypocalcemia (see page 380)
 - 3 Nutritional disturbances e.g. hypoproteinemia (see page 58) and deficiencies (see pages 60-64)
 - 4 Psychogenic disturbances e.g. fixation on GI tract or anxiety regarding sphincter mishaps in cases of long standing diarrhea

C Diet

- 1 Non irritant foods Many clinicians feel that food should be withheld or that the intake during the first 24 hours should be restricted to liquid foods. (See bacillary dysentery page 276) During the acute phase of enteritis the only foods which should be taken by mouth are the following: very bland items: water, weak tea, rice or barley gruel, meat broth, precooked cereals, toast, butter and soda crackers with butter and soft cooked (not fried) eggs. These food are usually administered in about that same order as tolerated.
- 2 Bland foods (never highly spiced or seasoned) These foods should be incorporated in the diets of patients convalescing from acute diarrhea or with chronic diarrhea. They include in addition to the nonirritant food the following items: cereals with milk or cream, thin broths and soup, bland cheeses, fish, fowl and meat (not fried), potatoes (not fried), breads, milk products, eggs and food beverages (not carbonated).
- 3 *Avoid Vegetables and fruits (especially raw), fried foods, bran, whole grain cereals, jams, jellies, preserves, syrups and candies, pickles, relishes and spices, coffee, carbonated and alcoholic beverages.*
- 4 Supplemental vitamins The bland diet is a restricted diet and may further increase the vitamin deficiency induced by altered intestinal absorption. Patients with chronic diarrhea should probably receive vitamin in dosage comparable to those used for chronic vitamin deficiency states. Roughly this amount may vary from 4 to 10 times the normal maintenance dose (see page 58).

D Anti diarrheal Agents

- 1 Bismuth preparations These may be used for both acute or chronic diarrheas
 - a Bismuth Subcarbonate U.S.P. B.P. 1.2 Gm (15-30 gr) after liquid bowel movement or q.i.d.
 - b Bismuth Magma N.F. (bismuth hydroxide and sodium bismuthate) 4 cc (1 t.p.) after liquid bowel movement or q.i.d.

- c $\frac{3}{4}$ Bismuth subcarbonate 15 30 0 $\frac{1}{2}$ ss i
 Camphorated tincture of
 opium (paregoric) q s ad 120 0 $\frac{1}{2}$ iv
 Shake well
 Sig 4 cc (1 tsp) after liquid bowel movements or q i d
 d Milk of bismuth and par g ic (equal amounts of each)
 may b substituted for the above mixture using th same
 do e

- e $\frac{3}{4}$ B llaodona xtract 0 5 gr viias
 Bi muth sub arbonate
 Calcium lactate
 Kaolin 5X 30 0 3i
 Peppermint oil 2 drops gtt ii

Sig 4 cc (1 tsp) t i d a c and h s or after liquid
 bow l mov ments p r n (modified aft Bockus)

- 2 P ctin kaolin compounds Th se are availabl and are use
 ful mixtures Do e 15 30 cc ($\frac{1}{2}$ l oz) t i d a c and
 h s or after liquid bowel movem ts p r n
- 3 Albumin tannat This drug has been recomm ded as an
 adjunct to other m sures when dia charges are profus
 Dose 2 Gm (30 gr) t i d a c and h s o after liquid
 bowel mov ment p r n
- 4 Opi tes Opiates must be av ided in chronic diarrheas and
 ar pr fer bly v ided in acute dia hea unless th r i in
 tractable diar hea vomiting and colic Alway x lud th
 p sibility of acute surgi al abdominal di s e befor ad
 mini tering opi te
- a Campho ated Opium Tincture U S P B P (paregoric)
 (NOT Opium Tinctu e U S P) 4 B (1 2 dr) after
 liq id movem ts p r n o with bismuth (ee above)
- b Cod ine Ph ph t U S P 15 65 mg ($\frac{1}{4}$ l gr)
 a b ut after l quid b wel mo ements p r n
- 5 St ong opi tes Morphine and dihydrom rph n h uld b
 r s v d fo s lcted p t nts with ut s e di r h a
 who fault po d to m re o s rv ti m sures
- a M rphin S ifate U S P 8 15 mg ($\frac{1}{8}$ $\frac{1}{4}$ gr) sub t
 aft r liq id bowel movem nts p n Thi drug may
 p od nau ea and v miting
- b Dihyd omorphin Hyd hl id U S P (Dil wdie®)
 M y b ub tit ted for m rphin if th und i abl d
 ffect of morphin a e to be avoid d D 2 3 mg
 ($\frac{1}{32}$ $\frac{1}{20}$ g) i M aft l quid bow l mov m nt p r n
- 6 Antispasmodi and s dativ d ug (e pag 266) The anti
 pa modic dr g parti ularly wh n u d in combin tion with
 th barbit at s exert a favorable and mild antipe list iti
 t on It m y be n ary at tim to administer the v
 ious b llaodona b llaodona lik lkaloids t s p int of
 t i lity in o d r t a hiev th d ir d ff ct Antisp
 modi s d ti d g may be consid red the ag nts of ch e
 in th t tm t of hroni diarrh a ociat d with anxiety
 t n ion t t s
- 7 Dig stant dr g Hydro hloric id panc e tin, and bil
 salt at tim s gi d finite non p if relief Wh n there
 i demon t abl d si i ncy of th e ubstanc s repla em t
 th apy is mo triking (S sp ill disease s)

Treatment

There is no satisfactory treatment for carcinoma of the esophagus

- A Diet Soft or liquid food should be given as tolerated gastric feedings may be given in selected cases
- B Surgical Removal This is reserved for the few who have no demonstrable metastases and are good surgical risks
- C Deep Radiation Therapy This may be employed in selected cases when surgery is not feasible
- D Morphine sulfate or other suitable analgesic agents should be administered as necessary for relief of chronic pain, especially in terminal cases (see page 33)

CARDIOSPASM (code No 641)

Spasm of the lower end of the esophagus may be due to local or reflex causes. Dysphagia, epigastric pain, and regurgitation of undigested foods are the most common findings. X rays reveal dilatation above the site of obstruction.

Treatment

- A Soft or liquid food as tolerated
- B Autonomic drugs have been employed with variable and non-spectacular results. Large doses of parenteral antispasmodics are often ineffective. Recent experimental and clinical studies suggest that the sympatholytic agents may be effective (see page 266)
- C Mechanical dilatation of the cardia by graduated bougies may be necessary

DISEASES OF THE STOMACH AND DUODENUM

PEPTIC ULCER

(Gastric code No 640 951) (Duodenal code No 651 951)

An acute or chronic ulceration of any portion of the GI tract which may be exposed to the action of a digestive lesion may occur at any point in the lower esophagus, stomach, upper duodenum (most common), gastroenterostomy margins and in certain anomalous areas of the GI tract (e.g. Meckel's diverticulum). The ulceration may be simple or complicated by hemorrhage, perforation, scarring and obstruction or by malignancy.

Diagnosis

May be based upon

- A HISTORY
 - 1 Pain Classically there is postprandial (1 2 hours) or fasting epigastric discomfort, burning or pain usually relieved by bland food and/or alkalis
 - 2 Other symptoms Nausea, vomiting, flatulence, distention, hematemesis and melena
- B Physical Findings Often local tenderness in the epigastrium

- C Lab oratory Finding There may be
- 1 Abundant or excessive free HCl in gastric juice both with and without histamine injection
 - 2 Gross or occult blood in stools
- D X ray Evidence of Ulceration Based upon films and fluoroscopy of GI series Ulcer activity is usually indicated by presence of niche or irregularity of mucosal contour but sometimes evidence is indirect such as altered peristalsis pylorospasm gastric retention or persistent deformity. Repeat GI series may be necessary to demonstrate active ulceration in certain cases
- E Gastroscopic demonstration of ulcer crater

Diagnostic Criteria

- A It is not advisable to make a diagnosis of peptic ulcer unless there is or has been x ray or gastroscopic evidence of ulcer
- B In face of clear cut peptic ulcer history without laboratory confirmation it may be necessary at times to perform repeated GI series
- C Malignant disease should be suspected when the following findings are present
- 1 Location The lesion is located in the stomach particularly if in the prepyloric region high on the lesser curvature or if on the greater curvature
 - 2 Duration of symptom is short (no previous symptoms)
 - 3 Age of the patient is more than 40 years
 - 4 Failure of response There is failure of clinical and x ray response after not less than 3 and not more than 4 weeks of intensive medical therapy (see below)

ACUTE PHASE

Treatment

A General Measure

- 1 Rest (physical and mental) The patient should have 2 or 3 weeks rest from work if possible. If the home situation is unsatisfactory or unpleasant or if cooperation of the patient is unsatisfactory hospitalization is recommended. If patient's finances are limited it may be necessary for him to continue work during treatment. In such case it is essential that he be given careful instructions regarding the carrying out of the medical program under the given working conditions. When possible arrangement should be made for a stepwise increase of his sleep and for any other factors which need to be modified in the patient's home or working environment.
- 2 Orientation The patient should be advised as to the chronic current nature as well as to the potentialities of the disease. Do not emphasize cancer as a complication of the disease.
- 3 Psychotherapy Anxiety producing mechanism should be relieved whenever possible. It is not usually wise to institute active psychotherapy during the acute phase of the illness (see page 36).
- 4 Alcohol and tobacco must be avoided.

- 5 Avoidance of certain drugs (e.g. corticotropin (ACTH) and the cortisones). Recrudescence of symptoms and even perforation and hemorrhage may occur in patients with peptic ulcer during the course of administration of corticotropin (ACTH) and the cortisones. This hazard should always be considered. The mechanism is not known.

B Diet

- 1 A wide variety of diets are available but the 5 popular diets or modifications of these are probably the most effective (see page 54). The patient should learn the principles of his diet and should be taught to be careful of his diet for the remainder of his life. Rich, spicy, hot and coarse particle foods should be excluded from the diet permanently. Regular and frequent high protein meals, with proper mental attitude during meals, should be emphasized as essential for successful results from diet. The length of time the patient should remain on each phase of the diet will depend on numerous factors, namely:
 - a Severity of symptoms
 - b Treatment situation (e.g. Sippy diet does not meet the nutritional requirements of the hard laborer. Additional food is essential)
 - c Intelligence and cooperativeness of the patient
 - d Response to treatment
- 2 Avoid short cuts. In general, most of the short cut or so-called modified methods do not ultimately save the patient time. In many cases they not only fail to provide the necessary relief of symptoms but also actually serve to lengthen the period of convalescence. The psychological importance of a strict dietary regimen in the acute phase is of great importance to peptic ulcer patients, especially to new patients. Even these patients will otherwise fail to recognize the importance of diet in the long term care of their disease. Patients on short cut diets become sloppy and lackadaisical and indifferent to the potentially serious nature of their illness. Unfortunately, there is no unanimity of opinion among clinicians regarding this matter of diet in peptic ulcer.
- 3 Protein hydrolysate solutions or similar commercial protein preparations may be used to supplement the Sippy diet especially when it is necessary to reduce the fat intake. Boiled cow's milk, protein separation, or goat's milk may be substituted in individuals intolerant of cow's milk.
- 4 Restrictions. Meat extract, bran, raw vegetables and fruits, fried foods, condiments, spices, alcohol, coffee, tea, and carbonated beverages.

C Drugs

- 1 Antacids. It is difficult to state which of the many antacids available are most effective since no certain clinical comparison of the agents listed below is possible. Some clinicians feel that antacids are of little or no value in the well-managed patient. In general, however, a suitable dietary regimen, disappearance of anticholinergic drugs will either obviate or decrease the need for antacids. All patients on antacid therapy should be watched for

diarrhea constipation alkalosis and fecal impaction

Antacid powders are prescribed on various schedules according to the stage of the diet. During the early stage of the Sippy regimen the powders are given on alternate hours or half hours during the day and even at night if necessary. The interval between powders may then be lengthened according to clinical and x-ray evidence of improvement. For more prolonged use the powder are usually administered 2 hours p.c. and p.r.n. Magnesium oxide is a laxative drug and calcium carbonate tends to produce constipation.

a Magnesium Oxide U.S.P. B.P. and Calcium Carbonate U.S.P. B.P.

R Magnesium oxide 30.60 0 Si i

Calcium carbonate q.s. ad 120 0 Si v

Sig. Take $\frac{1}{2}$ t. tsp. in half glass of water as directed. By varying the magnesium oxide in the above powder the laxative or constipating effects of the 2 ingredients may be effectually balanced. The powder may be given in alternate doses with the aluminum hydroxide gels (see below).

b M Bi Cal (magnesium bismuth aluminum) mixture

R Magnesium oxide 20.60 0 3v ii

Bismuth subcarbonate 20 0 3v

Calcium carbonate q.s. ad 120 0 3v

Sig. $\frac{1}{2}$ t. tsp. in half glass of water as directed. The bismuth incorporated because of its soothing coating effect this powder occasionally offers relief when the magnesium oxide-calcium carbonate powder fails to relieve.

Magnesium Trisilicate U.S.P. B.P. $\frac{1}{2}$ t. tsp. in half glass of water as directed. An excellent non-systemic antacid with unusual protective properties.

d Aluminum Hydroxide Gel U.S.P. (Amphogel[®], Creamalin[®], etc.) The emulsions have recently enjoyed popular use because of convenience of administration rather than for maintenance of alkalinity and because of its adsorbent protective and demulcent action. However, they occasionally interfere with phosphorus and vitamin absorption and may require large doses and occasionally fail to give relief.

(1) Aluminum hydroxide gel liquid 1 t. tsp. in half glass of water every 2-4 hours p.r.n.

(2) Aluminum hydroxide gel (d) tablets chew 1-2 tablets and follow with half glass of water every 2-4 hours p.r.n. The tablets are especially convenient for patients who are fitted to continue work or to travel.

(3) Aluminum hydroxide gel-magnesium trisilicate mixtures liquid (Gelusil[®], Trireamlat[®], etc.) 1-2 tsp. in half glass of water every 2-4 hours p.r.n. The addition of magnesium trisilicate increases the neutralizing power and protective coating action of the aluminum hydroxide gel. This mixture is also less constipating.

- (4) Aluminum hydroxide gel (dried) magnesium trisilicate tablets chew 1 or 2 tablets every 2-4 hours p r n and follow with a half glass of water
- 2 Sedative drugs The use of sedative drugs will depend on the emotional status of the patient. Tense and apprehensive patients will usually profit greatly from proper sedation. Most patients with peptic ulcer profit by sedative drugs. The barbiturates are the preferred sedatives. They may be used alone or in combination with antispasmodic drugs. Hypnotic doses of the barbiturates may be necessary to insure sleep during the acute phase of the ulcer (see page 39)
- 3 Antispasmodic antisecretory drugs
- a Belladonna preparations when employed in proper doses are probably as effective as any of the many new anticholinergic preparations and have the added advantage of being inexpensive
- (1) Belladonna Tincture U S P B P 0.3-0.8 cc (5-10 drops) in half a glass of water orally t i d 20-30 minutes a c and h s p r n (0.6 cc of the tincture equals about 0.2 mg of atropine). This preparation permits rather delicate titration of desired antispasmodic effect by simply regulating the number of drops
- (2) Belladonna Extract U S P B P 5-15 mg ($\frac{1}{8}$ - $\frac{1}{4}$ gr) tablets or capsules orally t i d 20-30 minutes a c and h s p r n (15 mg equals about 0.2 mg atropine alkaloid)
- (3) \mathcal{R} Belladonna tincture 10-30 0 (3i ss 3i)
Elixir of phenobarbital q s ad 120 0 (3iv)
Sig 1 t p in half glass of water t i d 20-30 minutes a c and h s p r n
- (4) \mathcal{R} Belladonna extract 5-15 mg (gr $\frac{1}{8}$ - $\frac{1}{4}$)
Phenobarbital 15 mg (gr $\frac{1}{4}$)
Sig 1 tablet t i d 20-30 minutes a c and h s p r n
- b Anticholinergic antispasmodic drugs Many new and effective but relatively expensive agents are available. These drugs as is the case with belladonna are given t i d q i d and should generally be given in doses large enough to produce some oral dryness. Although they are said to be relatively free of side effects it is best to watch for blurred vision difficulty in chewing constipation and urinary retention
- (1) Adiphenin hydrochloride (Trisentine[®]) 75 mg ($1\frac{1}{4}$ gr)
- (2) Methantheline Bromide U S P (Banthin[®]) 50-100 mg ($\frac{3}{4}$ - $1\frac{1}{2}$ gr)
- (3) Propantheline Bromide N N D (Pro-Banthine[®]) 15-30 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr)
- (4) Oxyphenonium Bromide N N D (Anirhyl[®]) 5-10 mg ($\frac{1}{12}$ - $\frac{1}{6}$ gr)
- (5) Diphenamid Methylsulfate N N D (Prantal[®]) 100-200 mg ($1\frac{1}{2}$ -3 gr)
- (6) Methacopolamine Bromide N N D (Pamine[®]) 2-5 5 mg ($\frac{1}{4}$ - $\frac{1}{12}$ gr)

CONVALESCENT PHASE

Treatment

- A. Re-examination.** When clinical quiescence of the lesion is evident (based on freedom from symptom) a repeat GI x-ray series is advisable to determine whether or not the evidence of healing of the ulcer is the case. In the case of gastric lesions failure of clinical improvement and x-ray improvement of the ulcer crater within a period of 3-4 weeks on a careful medical regimen should be taken as suggestive evidence of gastric malignancy.
- B. Education of Patient Regarding Recurrence.**
1. The patient should be educated to an understanding of the chronicity and recurrent potentialities of his ailment as well as of the dangers due to complications which may follow a neglectful or improper treatment.
 2. Factors causing recurrence. It should be emphasized to the patient that the following factors are most frequently responsible for recurrence of ulcer:
 - a. Improper diet and irregular eating habit
 - b. Irrregular living habit: long or irregular hours
 - c. Use of alcohol or tobacco
 - d. Emotional distress
 - e. Infections particularly of the upper respiratory tract
 3. Management. Patients should be instructed to return to the 3 ppy regime or to a modification thereof in the event of recurrence of symptoms or even prophylactically if they are exposed to conditions known to aggravate peptic ulcer. In addition to diet information antacid and other medication should be readily available to the patient.
- C. Rest and Relaxation.** Provision should be made for proper rest, eating vacation and the like to promote physical and mental relaxation.
- D. Psychotherapy.** Selected patients should be considered for individual or prolonged psychotherapy.

TREATMENT OF COMPLICATIONS

INTRACTABILITY TO TREATMENT

Although numerous cases undoubtedly exist where benign peptic ulcer fails to heal despite optimum medical upervision it is probable that most are cases of stubborn and persistent ulceration of an intractable medical regime or of partial or complete failure of cooperation on the part of the patient. The factors previously mentioned as being responsible for recurrence of peptic ulcer are often the same factors which interfere with the healing of ulcer. The diagnosis of intractability should be reserved only for those patients who have been given adequate and supervised trial of the 3 ppy. The possibility of malignancy or of other complications of the ulcer (e.g. pyloric obstruction, perforation, gastritis, etc.) must always be considered.

HEMORRHAGE

(Stomach code No 640 951 7) (Duodenum code No 651 951 7)

Although peptic ulceration accounts for about 70% of gross hemorrhage from the upper gastrointestinal tract one must consider the possibility of esophageal varices gastritis duodenitis carcinoma of the stomach hiatus hernia and systemic bleeding diseases

Treatment

A Emergency Measures for Hemorrhage and Shock

Refer to page 28 for general management of shock

- 1 Hospitalize patient at absolute bed rest
- 2 Warmth Keep patient comfortable If an ice bag is applied to the epigastrium avoid chilling the patient
- 3 Treatment of apprehension and anxiety
 - a Reassurance by word and manner of physician that the condition is not critical
 - b Rest Provide prompt mental and physical rest this can best be achieved in the hospital
 - c Sedation May be necessary
 - (1) Morphine should be avoided if possible since it may cause nausea Dose is 12-16 mg ($\frac{1}{2}$ - $\frac{1}{4}$ gr) subcut every 4-6 hours It is better to substitute codeine phosphate 30-65 mg ($\frac{1}{2}$ -1 gr) subcut or orally or dihydromorphinone (Dilaudid®) 4 mg ($\frac{1}{16}$ gr) subcut every 4-6 hours p r n
 - (2) Sodium phenobarbital (sodium phenobarbital) 0.03-0.1 Gm ($\frac{1}{2}$ - $\frac{1}{2}$ gr) subcut or orally during the first 24-48 hours
 - (3) Phenobarbital (phenobarbital) may be continued for several days if necessary
- 4 Oxygen Preferably by mask at 5-10 liters per minute (see page 145)
- 5 Transfusions There has been considerable controversy regarding the use of blood transfusions in bleeding ulcer However it is generally agreed that the previous conservative attitude (feeling that transfusion may raise the fallen blood pressure to a point at which occurrence of hemorrhage is warranted) Certainly in severe hemorrhage the time rate and volume of blood administration should suit the physiological needs and large amounts of blood may be given when indicated Transfusions must always be given if hemorrhage is severe (Hgb < 30% or RBC < 2.5 million) if immediate surgery is contemplated or if symptoms of anemia or shock are not rapidly controlled Slow and continuous administration of 500 to 2500 or more of whole blood daily may be necessary
- 6 Clinical and laboratory studies
 - a Take pulse respiration and blood pressure every 15-30 minutes this data may give information regarding shock status in advance of blood changes
 - b Observe all vomitus and stool for gross or occult blood
 - c Type and cross match the patient's blood as early as soon as possible Have whole blood or plasma available without delay

- d Obtain complete blood count and hematocrit initially and serially as indicated
- e Obtain blood N P N or urea nitrogen for comparison with late studies

B G I M E

1 Corrected hydration and salt depletion

Hypodermoclysis: Physiologic saline solution 1015 ml a daily by this method

- b Oral liquid feedings as soon as tolerated (see below)
- c Sodium chloride 3.6 Gm ($3/4$ 1 $\frac{1}{2}$ dr) may be added to each liter of liquid food mixture to prevent salt depletion

2 Nutrition

- a Starvation: The policy of initial starvation is subject to considerable controversy. Since the patient is often nauseated and anorectic, even in shock on the first day food may be safely withheld.
- b Fluids: If patient is nauseated or vomiting, thirst may be controlled by fluids given parenterally. The patient may be permitted to dissolve ice chips or hard fruit flavored candy under the tongue to relieve thirst.
- c Diet: If the patient is hungry and not vomiting, it is wise to begin immediately administration of bland food stuffs.

(1) Liquid diet: It is best to begin with a liquid diet of bouillabaisse of milk and cream mixture (see page 52). Increasingly, upper intestinal antacid powders. There to 6 Gm ($3/4$ 1 $\frac{1}{2}$ dr) of sodium chloride may be added to each quart of milk cream mixture to prevent salt depletion.

(2) Solid bland foods

- (a) Conservative approach: Solid bland food may be added when the patient has shown appreciable clinical improvement on the liquid (milk and cream) regime within 12 weeks and when the patient's stools have shown no occult blood for 2-3 days.
- (b) Liberal approach (e.g. Mullen's acid): This method permits immediate feeding of all non-irritant high caloric foods but in purified form.

C Con val sc i Care: Following the acute episode the conservative regimen such as outlined for uncomplicated peptic ulcer (see page 263) should be instituted.

D S u rg e r y: Surgery should be considered if

- 1 The general condition of the patient fails to improve despite the above measures.
- 2 Bleeding persists as evidenced by gross or occult blood in stools if the patient's condition permits a gastrointestinal x-ray. It should be performed to help localize the source, identify the character of the bleeding lesion. Manipulation during such examinations should be as gentle as possible. If ophogastrectomy are eliminated as a cause of bleeding and the bleeding persists for more than 2-3 weeks, perhaps the patient promptly for surgical intervention. Do not wait until the patient becomes a poor operative risk before making this decision.

PYLORIC OBSTRUCTION (code No 618)

It is important to differentiate pyloric obstruction due to spasm and edema from that due to scarring. The former condition may respond to medical treatment whereas the obstruction due to scarring is a surgical problem.

Treatment

- A Medical Measures (for obstruction due to spasm or edema)
 - 1 Bed rest preferably in the hospital
 - 2 Liberal use of antispasmodics (see page 266)
 - a Oral If patient is able to retain oral medication
 - (1) Tincture of belladonna 10-20 drops t i d or q i d
 - or (2) Belladonna extract 15 mg ($\frac{1}{4}$ gr) t i d or q i d
 - b Parenteral If the patient is unable to retain medication by mouth atropine sulfate 0.5-0.6 mg ($\frac{1}{200}$ - $\frac{1}{100}$ gr) t i d or q i d subcutaneously
 - 3 Sedatives
 - a Phenobarbital (phen barbitone) 15-30 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr) t i d q i d
 - or b Phenobarbital sodium (phenobarbitone sodium) 0.665 Gm (1 gr) subcut every 8-12 hours p r n
 - 4 Nutrition
 - a Sippy I diet should be used initially gradually progressing to Sippy II, III and IV as tolerated (see page 54)
 - b Fluid or mineral imbalance must be corrected if vomiting is severe or prolonged Parenteral method are most satisfactory (see page 57 and 74)
 - c Hypoproteinemia must be corrected since the resultant edema may increase pylorospasm
 - 5 Control of hyperacidity
 - a Gastric sections should be aspirated every morning and night with a small gastric tube. Some clinicians feel that continuous gastric suction should be employed initially
 - b Antacids may be employed as for treatment of uncomplicated ulcer (see page 264) but avoid alkalosis from excessive use of soluble antacids since this increases pylorospasm
- B Surgical Measures (for obstruction due to scarring)
 - 1 Surgery is to be employed only when a thorough trial of conservative measures has failed
 - 2 The various recommended surgical procedures will not be discussed. It is currently the practice to perform gastric resection in most cases although some surgeons favor gastroenterostomy

PERFORATION DUE TO ULCER

(Stomach code No 640 951 3) (Duodenum code No 651 951 3)

Acute subacute and chronic perforation of peptic ulcers may occur. Acute perforation constitutes a medical emergency. Immediate surgical repair, preferably by simple surgical closure, is indicated. More extensive operations are generally unwise at the time of the acute episode because of the increased operative hazard due to the patient's usually poor physical condition. If the patient has been receiving corticotropin (ACTH) or cortisone, these drugs must be discontinued. If the patient has had no previous therapy or if previous therapy has been inadequate, he may then be placed upon a conservative medical regimen. If the patient has had an inadequate trial of therapy prior to the episode, prepare him for possible further extensive operative procedures by transfusions and other supportive measures. The treatment of subacute or chronic perforation may be medical or surgical, depending upon the presence or absence of complications (e.g., abscess, involvement of neighboring viscera) or upon the persistence and severity of symptoms.

GASTRITIS (code No 640 3)

The stomach may become acutely or chronically inflamed due to a wide variety of specific and nonspecific causes. The symptoms of gastritis are poorly defined and variable. It is not unusual to have an absence of symptoms when present they loosely resemble the symptoms of peptic ulcer.

Physical examination is usually not diagnostic. X-ray findings are not remarkable unless the gastritis is complicated by atrophic hypertrophic erosive or sclerosing chronic changes.

Gastric acidity may be decreased in the chronic forms of gastritis and blood may be observed in the gastric contents and in the feces. Gastroscopic visualization may reveal characteristic mucosal changes which form the basis for clinical classification of gastritis. Patients with pernicious anemia who have atrophic gastritis must be observed carefully and periodically for evidence of malignant degeneration of polyps.

Treatment

- A General Measures. A dietary and drug treatment program essentially similar to that employed in peptic ulcer is useful (see page 264).
- B Specific Measures. Remove or eliminate specific causative and aggravating factors (e.g., infection, alcohol, tobacco).

GASTRIC MALIGNANCY

(Carcinoma of the Stomach) (code No 640 8)

Carcinoma of the stomach should be suspected in all patients over 45 years of age who develop dyspepsia. This disease occurs more commonly in men than in women. Lesions occurring in the region of the greater curvature and prepyloric area are usually malignant. A high index of suspicion, careful x-ray gastroscopic

272 Diaphragmatic Hernia

studies and gastric analysis afford the greatest opportunity for early diagnosis. Unfortunately by the time the disease is manifest metastases usually have occurred and the lesion is no longer amenable to satisfactory surgical therapy.

Treatment

- A Specific Treatment (corrective). Early and thorough gastric resection is essential if the patient is a good operative risk. Patients should be afforded the opportunity of corrective surgery regardless of the apparently advanced nature of a malignant lesion.
- B General Measures (palliative). To be considered only when corrective surgery is impossible.
- 1 Simple shunting procedures (e.g. gastroenterostomy) in the event of pyloric obstruction.
 - 2 Symptomatic and supportive treatment as indicated.
 - 3 Narcotics should be given in adequate doses to alleviate suffering (see page 23).

DIAPHRAGMATIC HERNIA

(Congenital code No 275 037 9) (Traumatic code No 274-424)

Herniation of a portion of the abdominal viscera through a congenital or aquired defect (especially esophageal hiatus) of the diaphragm may be manifested by a wide variety of symptoms but classically by epigastric distress and dyspnoea noted especially on lying down after meals. Nausea vomiting, small haemorrhage and angina-like symptoms may occur. X-ray demonstration of the hernia is usually necessary to confirm the diagnosis. Small esophageal hiatus hernias which are of questionable clinical significance are reported frequently (at least 10%) on routine x-ray gastrointestinal series.

Treatment

- A Treatment of reflux and dyspnoea (see page 260)
- 1 Small frequent feedings of bland easily tolerated food.
 - 2 Antispasmodic medication (see page 266).
 - 3 Antacid powders frequently provide relief from heartburn (see page 264).
- B Instructions to Patient
- 1 Patient should be instructed to:
 - a Avoid lying down immediately after eating.
 - b Avoid exerting vigorously after eating.
 - 2 Patient should be advised to sleep in the semi-Fowler position, or at least with upper part of body slightly elevated in an attempt to decrease acid regurgitation into the esophagus.
- C Surgical Treatment. 1 Correction of the hiatal defect is necessary if the following procedure and hospitalization are indicated only if the symptoms are progressively increasing and fail to respond to conservative medical management.

DISEASES OF THE INTESTINES

REGIONAL ILEITIS (code No 644 952)

An acute or chronic inflammation of the distal portion of the small intestine characterized by ulceration and scarring and often associated with internal and external fistulae. The condition must be differentiated from other specific causes of enterocolitis (gastrointestinal tuberculosis, chronic bacillary and amebic dysentery). The history often of long duration, is one of mild intermittent diarrhea and abdominal cramps relieved by bowel movement. The acute form may simulate appendicitis.

Physical findings may include tender masses in right or left lower quadrants, fistulous tracts and perirectal abscesses. Occult blood is often present in the white stools. Gastrointestinal x-ray series (small bowel study) reveals a loss of the mucosal pattern with narrowing and irregularity of the terminal ileum (string sign).

Treatment

A Corrective. Radical primary resection of the involved portion of the bowel is the procedure of choice after a seasonable period of conservative medical therapy has been tried. Despite extensive surgical treatment the disease will recur fairly often.

B Palpative

- 1 Diet. Bland high calor high vitamin adequate in protein.
- 2 Symptomatic treatment of anemia, diarrhea, avitaminosis as indicated.
- 3 Sulfonamides and antibiotics. Although of doubtful value, the sulfonamides which are poorly absorbed from the gastrointestinal tract might be given till (page 500). The effect of neomycin and chlortetracycline (Aureomycin®) have not been completely evaluated.
- 4 Corticotropin (ACTH) and corticosteroids may produce beneficial results in certain patients with regional enteritis but results have been quite variable and generally not too encouraging. Experience would indicate that long term use of these agents may not be without hazard and may result in increased destruction of intestinal tissue.
- 5 Palliative surgery. Short circuiting operation may be necessary when involvement is extensive and complicated.

DIVERTICULOSIS (code No 660 642)

DIVERTICULITIS (code No 660 642 0)

Multiple acquired or congenital outpocketings (pouch-like projections) may occur at any place along the course of the bowel, especially the colon. In diverticulosis the lesions are asymptomatic and are discovered accidentally on x-ray examination. Inflammation of diverticula (diverticulitis) with symptoms of intestinal abdominal inflammation referable to the involved site occurs mainly in individuals above 40 years of age. Variable lower gastrointestinal symptoms occur depending on location of diverticula. Abdominal pain and tenderness of rectum and bowel disturbance, quieted after

entiation from acute appendicitis. Laboratory evidence of inflammation may exist and x ray demonstration of diverticula helps to confirm the diagnosis.

Treatment

- A General Measures Conservative medical treatment is the method of choice.
- 1 Bland diet as tolerated
 - 2 Anticonstipation measures (see page 254)
 - 3 Coating agents. Non constipating antacid coating powders and gels, vegetable oils (olive oil), mineral oil or vegetable gum laxatives may be used (see page 255)
- B Surgical Treatment May be indicated in the event of complication.

ULCERATIVE COLITIS (Nonspecific) (code No 860.951)

An acute or chronic inflammatory disease of the colon of undetermined cause. It responds poorly to treatment and has a strong tendency to chronicity and relapses. Symptoms include (1) disturbance of lower bowel function with either diarrhea (usual) or constipation and with blood or bloody mucus in the stools and (2) systemic disturbances, anorexia, weight loss, and then fever, anemia, and even profound debility.

Proctoscopic examination reveals a diffusely superficially ulcerated bleeding mucosa. The presence of specific infectious organisms should always be ruled out by agglutination tests, stool examination for ova and parasites, and stool culture. Barium enema x ray appearance is that of decreased colonic caliber, loss of haustral marking, and indistinct outline of the colonic mucosa.

Treatment

A General Measures

- 1 Rest. Bed rest is advisable, usually only in the acute phase. Adequate rest periods can form an effective part of the daily routine of most patients.
- 2 Nutrition
 - a Diet. The diet should be bland, yet as appetizing and nutritious as possible. A bland, high calorie, high protein, high vitamin diet is recommended. When anorexia is marked, it is permissible at times to use other than bland foods if the patient so desires. These patients can often tolerate meat fairly well.
 - b Elimination diet. If allergic factors are suspected, elimination diets may be employed to aid diagnosis.
 - c Supplemental vitamins may be administered especially if nutrition is markedly disturbed.
 - d Ferrous sulfate 0.2-0.4 Gm (3-6 gr) tid po may be used to combat hypochromic anemia.
- 3 Psychotherapy. The exact role of psychological factors has not been determined. Certain personality type apparently are predisposed to the illness. In any case, anxiety-producing mechanisms should be eliminated where possible (see page 367). These patients need consideration and understanding and reassurance.

4 Symptomatic and supportive treatment

- a Diarrhea The various antiperistaltic agents employed for any chronic diarrhea may be used (see page 258). Use of narcotic drug should be avoided if possible except for severe acute diarrhea. Antispasmodic drugs are often of value (see page 266). Metamucil® or other vegetable mucoilage have been suggested to increase the bulk of the stool.
- b Nausea and vomiting (see page 251)
- c Cramps or tenesmus Hot water bottle to the abdomen and small antispasmodic medication may be employed.
- d Hypoproteinemia This is best corrected by high protein diet when restrictions are well tolerated. Protein hydrolysates dissolved in acceptable liquids such as milk, fruit juice, broth, etc. may be used to supplement the dietary protein when indicated. If hypoproteinemia is marked and cannot be corrected by dietary means, blood plasma or whole blood transfusion may be necessary.
- e Bleeding tendency (due to hypoproteinemia) Treatment with menadione or other vitamin K preparations (see page 81 and 228) may be indicated.
- f Nervousness In addition to general psychotherapy, measures mentioned above, mild sedation is often necessary. These patients often show little response to various antispasmodic sedative mixtures (see page 266).

5 Antiinfective agents These should not be considered to be specific for curative in the diseases. However, good results and prolongation of remissions have been reported by some observers.

Sulfonamides Many preparations have been used. Isoniazid, sulfonamide, which are poorly absorbed from the gastrointestinal tract, are preferred. Sulfisoxazole, USP (Sulfisoxazole®) and Phthalylsulfathiazole, USP (Sulfathiazole®) are more enjoyable to tolerate than the others. Observe the usual precautions for sulfonamides (see page 500).

- (1) Sulfisoxazole® 3.0 Gm (45 gr) very 4 hours
(Should not be used if the patient has diarrhea)
- (2) Sulfathiazole® 1.5 Gm (22½ gr) every 4 hours
(This dose may be doubled in cases of severe disease)

- b Antibiotics The tetracycline drugs have been employed but are of questionable value. Oral penicillin is of little value. Chlorotetracycline, USP (Achromycin®) is preferred on the basis of limited experience to be effective when given in doses of 2.4 Gm daily for 10 days. Doses of 4.6 weeks. Oxytetracycline, USP (Tetracycline®) and Chloramphenicol, USP (Chloromycetin®) are probably about as effective as the tetracycline.

6 Corticosteroids (ACTH) and the corticosteroids are known to induce remissions in many cases of ulcerative colitis. They are usually effective but may induce temporary suppression of the inflammatory exudative process. Their use may be considered in the circumstances when clinical control is difficult and the activity of the disease is interfering with other treatment measures.

Optimum dosage schedules have not been established for these drugs. They are usually given in courses during exacerbations. They should be administered in high doses and gradually reduced as symptoms disappear. (For dosage see page 423.) Reports indicate that relapse recurs when the drugs are discontinued before onset of the natural cyclical remission phase. It seems the drugs are used most profitably as adjuvants in the control of acute exacerbations of the disease and should probably be avoided for long term use.

- B Surgical Measures** These should be used only after careful prolonged medical therapy fails. This does not mean that the patient should be terminal or a poor operative risk before operation is considered. The choice of surgical procedures will not be discussed here. Subtotal or total colectomy may be employed initially or resorted to only after an extended trial of ileostomy or colostomy.

BACILLARY DYSENTERY (code No 660 116)

Bacillary dysentery is an acute or chronic infection with the dysentery (*Shigella*) group of bacilli. Involving primarily the colon. It is characterized by diarrhea, abdominal cramp, tenesmus and sigmoid tenderness. Stools contain blood, pus, mucus and microscopically large non-motile microphages (coliforms). Stool cultures are positive during the first week; agglutination tests are positive during the second week. The important bacteriological and clinical types include:

Organism	Distribution	Severity	Significant Agglutination	Fermentative	
				Lactose	Mannitol
<i>Shigella</i> <i>flexneri</i>	Tropics and subtropics (epidemic)	Severe	< 40		
<i>Shigella</i> <i>flexneri</i>	World wide	Mild	> 160		+
<i>Shigella</i> <i>sonnei</i>			< 320	+	+

Treatment

A Emergency Measures

1. Isolate patient and use all contagion precautions.
2. Overcome dehydration and electrolyte imbalance by the liberal use of saline and dextrose solution. I.V. and when necessary the use of potassium solutions (see page 22). Urinary output should be kept at 1000-1500 cc (1 1/2 qt) per day.
3. Watch for circulatory collapse and shock in severe case (see page 27).
4. Stool examination. Obtain specimen for microscopic examination and culture to determine causative organisms.

B Specific Measures

The sulfonamides have been considered effective in the treatment of bacillary dysentery but the

broad spectrum antibiotic appear to be gaining in favor. There is a significant variation in response of specific organisms in different individuals. Sulfur treatment is probably indicated only in severe acute Shigella infections and is used to combat the toxin of the infection.

1. **Antibiotics.** Administer tetracycline drugs and chloramphenicol in doses of 0.25 to 1.0 Gm every 6 hours.
2. **Sulfonamides.** Observe the usual precautions for sulfonamides and watch for signs of possible toxicity. Sulfadiazine is considered the drug of choice. Give 2.4 Gm (30-60 gr) Stat with equal or double quantities of sodium bicarbonate and follow with 1.2 Gm (15-30 gr) every 4 hours per os. If diarrhea is severe, larger doses by mouth or the use of sodium sulfadiazine parenterally may be necessary.
3. **Sulfur treatment.** Administer sulfonamide or broad spectrum antibiotics simultaneously.
 - a. **Bacillary dysentery polyvalent antitoxin serum** (not antibacterial). Test for sensitivity and administer 30-100 cc diluted tenfold in a saline solution I.V. tid until the toxemia is over.
 - b. **Shiga antitoxin serum.** Administer as above in doses of 40-80 cc in 50% of physiologic saline solution I.V. bid.

C. General Measures.

1. **Isolation.** Disinfect all body discharges completely in and about the patient.
2. **Bed rest.** When diarrhea is severe and patient is weak, it may be advisable to have the patient discontinue all ambulation, also bent position, to avoid the physical exertion necessitated by use of the bedpan.
3. **Initial purgation.** Control rectal and probably not advisable.
4. **Careful rectal hygiene.**
5. **Local heat treatment.** abdominal pain.
6. **Sedation.**
 - a. **Phenobarbital** (phenobarbitone) 15-30 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr) orally tid or qid.
 - b. **Pentobarbital sodium** (pentobarbital sodium) 0.1-0.15 Gm ($\frac{1}{12}$ - $\frac{1}{8}$ g) orally prn.
7. **Narcotics.**
 - a. **Codeine Phosphate U.S.P.** 15-65 mg ($\frac{1}{4}$ -1 gr) orally or about prn pain.
 - b. **Compound Tincture of Opium U.S.P.** (paregoric) 4-8 (1-2 dr) as necessary for pain and frequent loose bowel movements.
8. **Atropine Sulfate U.S.P.** 0.3-0.6 mg ($\frac{1}{200}$ - $\frac{1}{100}$ gr) orally subcutaneous for cramps.
9. **Fluids.** Adequate fluid intake by oral and parenteral routes should be forced to limit dehydration as indicated. Total oral fluid intake should approximate 3000 cc (3 qt) in the acute phase. On or more liter of parenteral saline solution daily may be necessary to replace fluid and salt loss in profuse diarrhea.

10 Nutrition *Avoid starvation diets* However in severe cases patients should not be allowed to eat a normal diet for 1½ to 2 months after the acute phase Give parenteral feedings if necessary Evaluate bowel symptoms before adding the various dietary constituents

- a Early acute stage Feedings of clear broths rice water albumin water tea with clove barley water or apple juice (not cider) at frequent intervals
- b Late acute stage Gradually add as tolerated boiled milk cereals and strained fruit juices toasted soda crackers or bread and gelatin desserts
- c Subacute stage Gradually add as tolerated mashed potatoes boiled rice boiled chicken soft cooked eggs lean fish steamed beef custards and puddings

Chronic Bacillary Dysentery

The clinical manifestations and treatment of chronic bacillary dysentery are similar to those of chronic ulcerative colitis (see page 274)

FOOD POISONING

The term food poisoning ordinarily refers to the acute intoxication which results from the noxious agents or enterotoxins produced by bacteria. This is in contrast to gastrointestinal disturbances which are actually the result of infection of the gastrointestinal tract with microbial organisms (see page 276) or which are due to poisons of vegetable animal or chemical origin (see pp 530 to 546). Food poisoning is a result of poor food hygiene either in preparation or in storage or distribution or handling. Suspicion of food poisoning should arise in instances of febrile gastrointestinal disturbance of acute onset especially when more than one individual in a family group or community is involved. A careful history and collection of specimens for a suspected food vomitus and stools for laboratory study may be indicated. Reporting to local health authorities is essential.

Treatment is symptomatic and supportive except in botulism for which specific antitoxins are indicated. Perform gastric lavage and withhold food sedation and parenteral fluids. Liquid and soft diets (see page 46) are indicated in convalescence.

Organism	Onset After Ingestion	Severity	Treatment
<i>Clostridium botulinum</i>	12-24 hours	Very severe often fatal	Antitoxin (see page 473) Symptomatic and supportive
<i>Staphylococcus aureus</i>	1-6 hours	May be usually recover in 1-4 days	Symptomatic and supportive
<i>Salmonella enteritidis</i>	8-24 hours	May be usually recover in 1-2 days	(see pp 252 and 258)
<i>Streptococcus faecalis</i>	5-20 hours		

HEMORRHOIDS (Piles)

(Internal code No 66x 641) (External code No 67x 641)

Treatment

Treatment is directed at the 3 commonst disturbing manifestations bleeding prolapse and pruritus

A General Measures

- 1 Control constipation (see pag 254) Mineral oil 15-30 cc (1/2 to 1 oz) h a and in early a m may be especially effective
- 2 Use of soft toilet tissue slightly moistened with water and good rectal hygiene are essential

B Specific Measures1 Bleedinga Palliative measures

(1) Local astringent suppositories

() Tannic acid suppositories 0.2-0.3 Gm (3-5 gr)

(b) Bismuth subgallate suppositories 0.3-0.5 Gm

(5-7 1/2 gr)

(2) Bed rest if necessary

b Surgical measures Injection or removal as indicated2 Prolapsea Palliative measures

(1) Replacement of prolapsed hemorrhoid with lubricated finger to prevent strangulation patient in lateral recumbent position

(2) Warm sitz baths 20-30 minutes t i d or q i d

(3) Local lubricant (petrolatum) to anal region

(4) Local anesthetic (e.g. Nupercaine® or benocaine) sedation and analgesia p r n

(5) Bed rest if possible

b Surgical measures as indicated3 Swelling and itching (see pag 80) Lubricants and local anesthetic ointment in addition to above general measures

CARCINOMA OF THE COLON (code No 660 8)

Carcinoma of the colon should always be suspected if patient is more than 40 years of age has had change in bowel habits rectal pain or bleeding or has unexplained anemia. Verify by digital examination (positive in 30% of cases) sigmoidoscopic examination (positive in 5%) and for lesions higher in the colon by barium enema

Early surgical resection may be curative. If the lesion is inoperable treat symptomatically for constipation anemia pain etc

DISEASES OF THE HEPATOBILIARY TRACT

INFECTIOUS HEPATITIS (code No 660 100)

An acute infectious disease due to an unknown filtrable agent. It is characterized by (1) prodromal symptoms of anorexia malaise nausea vomiting abdominal pain light fever and headache for 1-7 days (2) subsequent development of varying degrees of icterus

bilirubinuria for days or weeks and (3) convalescent period of weakness and easy fatigability for days or weeks. The epidemic type has an incubation period of 3-6 weeks. The homologous serum type usually has an incubation period of 3-4 months. Infectious hepatitis may terminate fatally.

Differentiate from influenza, infectious mononucleosis, malaria, cholecystitis, yellow fever, leptospirosis, drug intoxication, quinine (Atabrine®) discoloration, and carotemia.

Treatment

A. General Measures

1. Bed rest is absolutely necessary until the initial acute symptoms have subsided and should be maintained judiciously until there is no longer clinical or laboratory evidence of the acute disease. Absolute bed rest beyond the most acute phase is not warranted. The return to activity during the convalescent period should be gradual.
2. Nutrition. Keep a close check on patient's actual intake and output.
 - a. Fluids and liquid foods. If patient is unable to take or retain food or fluids by mouth.
 - (1) 5% glucose solution should be given I.V. with 5% protose in hydrolysate as needed to maintain nutrition and fluid balance. Saline infusions should be used only to correct unusual losses of chloride by vomiting, etc.
 - (2) Tub feedings of high carbohydrate formulas and skimmed milk as given below (and on page 59) should be used if vomiting is severe or protracted.
 - (3) The use of irradiated plasma has been recommended but it is doubtful if present irradiation methods are effective in viral sterilization. Parenteral administration of protein hydrolysate or salt pool albumin may be necessary to maintain nitrogen balance.
 - b. Restricted solid foods. When patient's appetite improves, small frequent oral feeding of any of the following may be instituted as tolerated.
 - (1) Fruit juices fortified with glucose every 2-3 hours.
 - or (2) Powdered skimmed milk in chilled water every 2-3 hours.
 - or (3)

Dextrose	6-8 Tbsp	(90-120 cc)
Milk	1 Quart	(960 cc)
 - (4) Supplemental vitamins, particularly of the B complex, may be incorporated to advantage in the feeding schedule if tolerated.
 - c. Full diet. As soon as the patient is able to eat an adequate diet, provide a high CHO and high protein diet (see page 58) and supplementary vitamins of the B complex.
 - d. Lipotrophic agents such as choline, yescin, or methionine are of doubtful value.
 - e. Avoid physical exertion, necessary transportation, alcohol, all medication, whenever possible, especially barbiturates, morphine, and sulfonamide, and surgery, especially with general anesthesia.
3. Corticotropin (ACTH) has been reported to apparently exert a favorable effect on the progress in certain cases of acute hepatitis. Experience has been limited.

hazards of corticotropin treatment in liver disease must be considered (see page 423)

B Prophylaxis

- 1 Isolation of infected individuals is recommended. Human immune globulin 10 c.c. 1 M. may prevent or ameliorate the disease if given to exposed subjects during the incubation period.
- 2 Avoid unnecessary transfusions especially of possibly infected blood serum or plasma.

CHRONIC HEPATIC DISEASE

(Laennec's Cirrhosis code No 680 956)

(Following Acute Degeneration code No 680 952)

Etiology

Industrial chemical hepatotoxic drug alcohol (in excess) viral hepatitis chronic infection chronic biliary tract disease chronic malnutrition (see also) congenital failure

Diagnosis

- A **Symptoms** Weakness sense of fatigability right upper quadrant discomfort variable symptoms of dyspepsia hematemesis melena abdominal swelling distended massive pruritus mental depression
- B **Physical Examination** Asthenia weight loss pallor spider angiomas peripheral edema icterus hepatomegaly hepatic tenderness ascites apudmascia collateral venous hemorrhoids dependent edema
- C **Laboratory Findings** Anemia icterus decreased bile function (as measured by multiple liver function tests) (see page 280) Unfortunately no single test and often no group of tests is entirely reliable
- D **Special Diagnostic Procedures** Liver biopsy (punch) esophagoscopy x-ray demonstration of esophageal varices

Treatment

A Specific Measures

- 1 Removal of exogenous aggravating agent
 - a Industrial and household toxins
 - b Alcohol and opiates
 - Drugs Coal tar drugs (e.g. sulfonamide fast acting barbiturates that are retarded by liver) anesthetic agents narcotics (e.g. morphine sulfate)
- 2 Removal of endogenous aggravating factors
 - a Thyrotoxicosis
 - b Urgent conditions Biliary tract disease pancreatic disease intestinal obstruction
 - Chronic infections Syphilis tuberculosis undulant fever amebiasis etc. Specifically cut viral hepatitis should be treated adequately to prevent chronic hepatitis in the future (see page 273)

B General Measures

- 1 Physical rest This is essential in the presence of hepatic ascites and jaundice

2 Adequate nutrition

- a Diet A high caloric diet (at least 2500-3000 Calories) is recommended. Carbohydrates should supply the principal caloric needs. Protein should be adequate but not too high especially in patients prone to protein intoxication. The diet should contain sufficient fat to be palatable. Patients with cirrhosis frequently complain of anorexia and so it is essential to make their diet attractive and palatable as well as highly nutritious. Tube feedings may be necessary. Fried or greasy foods, highly spiced foods, and alcoholic beverages should be avoided.
- b Vitamins A and D Absorption of vitamins A and D in liver disease is impaired due to biliary deficiency. Vitamin A 5000 units and vitamin D 1000 units 1-2 times daily may be given. Perles may be preferred when anorexia is present.
- c Vitamin B complex factors have been widely used but it is questionable that lipotropic or other actions of this group of vitamins exert any striking beneficial results except in those circumstances where there is a deficiency of these vitamins (due to inadequate food intake). It is felt that if vitamin B complex administration is indicated it is provided by the following:
 - (1) Dried Yeast U.S.P. (brewer's yeast) powder 15-30 Gm daily or 20-30 0.6 Gm tablets daily
 - (2) Vitamin B complex high potency preparations
 - (3) Crude liver extract 1-2 cc i.m. 1-2 times weekly
- d Amino acid supplements (Protein Hydrolysate N.N.D.) may be incorporated in oral tube or parenteral feedings as indicated:
 - (1) Oral 2-15 Tbsp t.i.d. (to supply 50-400 Gm daily)
 - (2) Tube 2-15 Tbsp t.i.d. Rule out varices first
 - (3) I.V. 5% solution with 5% dextrose 1-3 liters daily
- e Skimmed milk may be used for oral or tube feedings.
- f Salt poor albumin 50-100 Gm daily may be employed in severe cases (very expensive).
- g Ascitic fluid Readministration of ascitic fluid by sterile technic (if protein in ascitic fluid is greater than 1%).
- h Transfusions of whole blood if severe anemia and hypoproteinemia coexist.

3 Ascites and edema may be treated by

- a Low sodium diet Reduce sodium intake to less than 2 Gm NaCl daily (see p. 55) and even less if necessary. Diets severely restricted in sodium are apt to be nutritionally inadequate and unpalatable but remarkable improvement of ascites, edema, liver disease, and portal hypertension has been reported on patients who have been on diets containing 200 mg (3 gr) sodium per day for periods of a few weeks to more than two years. The danger of inducing the so-called low salt syndrome with renal failure and death must be considered and watched for.
- b Attempt to restore plasma proteins to normal levels (see above). This is very difficult to achieve.
- c Chlorothiazide (Diuril®) 0.5 Gm b.i.d. q.i.d. or lily is stated to be effective in producing diuresis in certain patients with cirrhosis with ascites producing a marked

- incr ase in the excretion of sodium p tas sum a d hlo ide Long range effect venous s de ff cts and potential toxic ity hav not been fully valuated
- d M rcorial d(uretics 1 2 cc 5 V or 1 M once or twice a week (see page 204)
- e Abdomin l paracente is for pain, discomfort or inability to at if neces ary
- 4 An mia
- a Hypochromic anemia Fe rous sulfate 0 2 0 3 Gm (3 4 1/2 gr) enteri coat di blets t i d p c
- b Hyperchromic macrocytic an mia Crud liver extract 1 2 cc 1 M on e or twice a week
- 5 H morrhagic tend n y du to hypoprothrombinemia may be treated with vitamin K preparation although this treatment is ineffectiv when intrahepat c damag is severe Blood transfusions m y be necess ry to cont ol the bleeding t nd ney Some c ution should be obs rved in using large doses of salicylat s in these patients becaus of the enhance d hypo-prothrombin effect
- a Oral M nadione U S P M naphthon B P 1 3 tablets of 1 0 mg (1/30 gr) each t i d p c If ob trusive jaundic is p esent give supplem ntary bile s lts (see page 286)
- b I V o 1 M M nadion Sod um Bisulfite U S P 2 mg (1/30 gr) ry other day
- 6 Hem rrh ge from esophag al varic s S vere bleeding can at times be controlled by the use of the triple lumn n tube Surgical m asures are usually ha ardous and unsatisfactory but surg ry to relieve portal hyp t on may be c sidered in selected patient In young r patients in oth rwise good cond ti n in wh m hepatocellular dysfunct on is r latly light porta av l an tomo is m y b of b nef t
- 7 Miscell neous problems
- a Pruritus (see p ge 66) n usea and vomiting (see pag 251) and constipation (p g 254)
- b Corti otropin (ACTH) and th rti on Th se ag nts should be us d w th careful on id r ti on of hass ds of h mor hag t ndency port l thromb ls and sodium r te ti on They are b st not u d in ad an d cirrhosis

ACUTE CHOLECYSTITIS (code No 587 100)

Acute inflamm t on of the gallbladder m y consi t of any one f wld variety f pathologi l ion of th gallbladd r which are difficult to diff r ntial clinically Th conditions may d velop a a r ult of obstruction of th bill y pa ag s with or without stones or as a result of inf ction Clinical finding may vary consid rably in individual ases but fo purp es of man g m nt th y may be con enic ly divid d into 3 groups a ording to sev lty mild interm di t and s v re

Di gn is

A Symptoms

- 1 Hist ry of chroni dyspepsia o p ov d biliary calculi may or m y not be ell ited

- 2 Attacks of right upper quadrant colic frequently nocturnal with residual gallbladder tenderness occur
- 3 Nausea and vomiting are usually present during acute episodes

B Physical Examination

- 1 Jaundice may occur during or following attacks
- 2 Localized right upper quadrant tenderness is common
- 3 Fever may be present or absent

C Laboratory Findings

- 1 Leukocytosis is inconstant
- 2 Icterus index is elevated in common duct obstruction
- 3 X ray findings are variable and at times difficult to interpret. A gallbladder which fills poorly with the dye or empties slowly may be normal. Demonstration of stone in the cholecystogram is the most important finding

Treatment

A Mild Type (Mild or indefinite symptoms. Doubtful diagnosis)

- 1 Bed rest. Make frequent observations including repeated gentle abdominal examinations as indicated
- 2 Routine laboratory studies and icterus index
- 3 I V fluids. If patient is vomiting administer 5% glucose in saline solution I V. Later when tolerated add bland oral food and fluids gradually to diet as in patients with a ulcer enteritis (see page 274)
- 4 Apply local heat or cold to the abdomen
- 5 Sedative or analgesic drugs as required (see below)
- 6 After a quiet episode has subsided carry out further diagnostic studies (e.g. GB series) as indicated
- 7 Elective surgery may be considered later if
 - a Episodes of acute cholecystitis are definite severe or recurrent
 - b Cholelithiasis especially if condition is symptomatic (or if asymptomatic and patient is less than 45 years of age)
 - c Secondary disease exists in related structures (e.g. liver, pancreas)

B Intermediate Type (Symptoms are moderate and definite and the patient's general condition is good but the subsequent clinical course is unpredictable)

- 1 Bed rest in hospital. Follow patient by careful frequent observations and noting of all signs and repeated gentle abdominal examinations as indicated
- 2 Perform WBC and differential count every 12 hours and icterus index or serum bilirubin daily (if jaundice is suspected or present). Blood electrolytes, NPN and serum amylase studies may be indicated
- 3 Plain x rays of the abdomen are of value if the patient's physical condition permits it. Postpone cholecystograms until acute phase has passed
- 4 Nasal duodenal suction should be employed for abdominal distention and vomiting
- 5 Fluid and electrolyte balance to be maintained by parenteral fluids as indicated
- 6 Sedation. Phenobarbital sodium (phenobarbitone sodium) 0.055 to 0.13 Gm (1 to 2 gr) or pentobarbital sodium (pe to

ba bitone sodium) 0.1 to 0.3 Gm ($1\frac{1}{2}$ to 5 gr) i. M. or by rectal suppository

- 7 Analgesia Use the following gly or in combination
 - a Codeine Phosphate U.S.P. 30 to 65 mg ($\frac{1}{2}$ to 1 gr) q 4 hours p.r.n.
 - b Morphine Sulfate U.S.P. 10 to 16 mg ($\frac{1}{8}$ to $\frac{1}{4}$ gr) q 4 hours p.r.n. as required only
 - c Atropine Sulfate U.S.P. 0.4 to 0.6 mg ($\frac{1}{150}$ to $\frac{1}{100}$ gr) q 4 hours p.r.n. for severe pain only
 - d Glyceryl Trinitrate U.S.P. (nitroglycerine) 0.3 to 0.6 mg (200 to 100 gr) under tongue p.r.n.

Papaverine 65 mg (1 gr) q 4 hours p.r.n.
- 8 Evaluation When clinical and laboratory evidence point to progression during the first day of observation or if they show no evidence of improvement after the first 2 days, operative intervention is usually indicated (cholecystectomy).
- 9 Elective surgery When there is clinical and laboratory evidence of improvement during the first day of observation the patient should be managed conservatively until symptoms have subsided.
 - a Perform necessary x-ray and laboratory studies to confirm diagnosis and to evaluate the patient's physiological status.
 - (1) Icterus index Surgical risk is less when this is falling than when it is rising.
 - (2) Choleystogram Preferably after jaundice clears.
 - (3) Liver function tests (see page 280) plasma prothrombin, serum proteins, and serum electrolytes may provide valuable information.
 - b Prepare for subsequent elective surgery by high CHO adequate protein low fat diet with supplemental vitamins, plasma, whole blood transfusion as needed and Menadione Sodium Bisulfite U.S.P. (vitamin K) 2 mg ($\frac{1}{50}$ gr) i.v. every other day if jaundice is present and prothrombin is low.
 - c Postoperative risk patients in the high and over 50 years of age and especially obese patients should be operated upon only after a full individual evaluation.
 - d Elective surgery Cholecystectomy should be performed only if the patient has been adequately prepared for operation. Common duct exploration is carried out at the time of cholecystectomy.
- C 5 C In cases of empyema, gangrene, perforation and bulging of the gallbladder, cholecystectomy is usually the treatment of choice. After cholecystectomy when the patient is a suitable condition has subsided, perform cholangiography to determine presence of stones and patency of common duct. If the patient is a good surgical risk, perform adequate follow-up operation (as above).

CHRONIC CHOLECYSTITIS (code No 587 100 0) CHOLELITHIASIS (Gallbladder) (code No 587-615)

As with acute cholecystitis chronic inflammations of the gall bladder may be grouped together clinically irrespective of the wide variety of pathologic lesions. The disease usually follows repeated attacks of acute cholecystitis or may be due to chronic biliary stasis, biliary stones or infection. Many patients who exhibit symptoms of so called chronic gallbladder disease are actually suffering from functional GI disturbances e.g. nervous dyspepsia (see page 260). The management of cholecystitis without stones is usually a medical problem. When associated with stones the condition often requires surgery. The question of surgery for asymptomatic cholelithiasis (silent gallstones) remains extremely controversial and the decision to operate must be individualized for each patient.

Diagnosis

- A History: Recurrent episodes of shifting upper abdominal distress, largely in the right upper quadrant, occasional episodes of acute colic, abdominal distention, nausea and vomiting and intolerance of fatty and gas forming foods.
- B Laboratory Examination
 - 1 Gallbladder dye demonstration of poorly functioning gall bladder (poor filling and emptying on repeated examination) and/or biliary calculi.
 - 2 Duodenal drainage may demonstrate excessive quantities of exfoliated epithelium, mucus, bacteria and pus in dark fraction of bile.

Treatment

A Medical Management

1 Indications

- a Patients without clinical or x-ray evidence of stones who respond to careful medical treatment.
- b Questionable diagnosis or low grade symptoms. Differentiate from functional dyspepsia (a difficult problem).
- c Patients who refuse surgical treatment.
- d Poor operative risk patients.
- e Patients with a short life expectancy from other cause.

2 Treatment

- a Diet: In general 2 different types of diet.
 - (1) Low fat diet (classical type). This diet excludes both cooked and uncooked fats from all sources (see page 54).
 - (2) No grease diet (modern concept). This diet excludes only the cooked fats (grease) which are non-emulsified at body temperature but includes the uncooked fats such as are emulsified at body temperature. The first phase of this diet is similar to the Sippy diet with frequent feedings of milk and cream as improvement occurs the diet incorporates egg, butter, cooked vegetables and fruit and cereals as tolerated.
- b Antispasmodic medication: Very useful.
 - (1) Tincture of belladonna 10 drops t.i.d. a.c.
 - (2) Belladonna extract 15 mg ($\frac{1}{4}$ gr) t.i.d. a.c.
 - (3) Phenobarbital antispasmodic mixtures (see page 268).

- (4) Atropine sulfate 0.4 to 0.6 mg ($\frac{1}{150}$ to $\frac{1}{100}$ gr) orally or sublingually or subcutaneously
- c Bile acid preparations Not to be used in patients with biliary fistula due to complete mechanical obstruction. A cholagogue stimulates evacuation of the gallbladder and choleric alters secretion of the bile constituents and hydrocholeretic alters volume of bile
- (1) Dihydrocholic Acid U.S.P. (Decholin®) 0.25 Gm ($\frac{3}{4}$ gr) tid p.c. choleric (?) hydrocholeretic
- (2) Ox bile Extract Capsules N.F. 0.3 Gm (5 gr) or tablets 0.2 Gm (3 gr) tid p.c. cholagogue choleric and hydrocholeretic
- d Sedation Phenobarbital antispasmodic mixtures (see page 268) and barbiturates (see page 39)
- e Antacids These drugs frequently provide empirical relief of many of the annoying symptoms of gallbladder dyspepsia. Their mode of action is not clear but they are felt to relieve associated hyperacidity and to have soothing effect on the duodenum and sphincter (see page 264)
- f Laxative drug (cathartic)
- (1) Sodium Phosphate N.F. (disodium phosphate) 4.8 Gm (12 dr or 12 tsp) dissolved in warm water before breakfast
- (2) Magesium Sulfate U.S.P. (Epsom salts) Dose 4.8 Gm (12 dr or 12 tsp) dissolved in warm water before bedtime. This may be used initially but its prolonged use is inadvisable
- g Local heat to abdomen Hot water bottle or electric pad preferred for mild discomfort

B. Surgical Management

- 1 Indications (providing the patient is a good surgical risk)
- a Patient with clinical or x-ray evidence of stone who fails to respond to intensive medical treatment. Surgical results however also questionable
- b Patients with biliary stones with or without jaundice who have persistent attacks of right upper abdominal quadrant pain. A symptomatic cholelithiasis in patients less than 45 years of age indicated by some to be an indication for surgery
- Patient with suspicion of gallbladder malignancy
- 2 Cholecystectomy is contraindicated in general cholecystitis if my preference is to the patient is poor except when the surgical risk is poor the patient is seriously ill then a tentative indication

DISEASES OF THE PANCREAS

ACUTE PANCREATITIS (code No. 690.930)

Acute pancreatitis is characterized by a sudden onset of severe agonizing constant or intermittent pain often extending to the mid back shoulder flanks. Symptoms of vasomotor collapse (shock) may be present. Paralytic ileus or obstipation and vomiting often occur. A pathologic study of dyspepsia ulcer and gallbladder diseases

may be elicited. Physical examination reveals epigastric tenderness and rigidity. There is usually abdominal distention. There is an elevation of serum amylase and lipase levels (may be transitory). Leukocytosis is common and glycosuria may occur.

Treatment

- A. Emergency Measures for Impending Shock** (Vasomotor Collapse) (see page 27)
1. Bed rest in shock position (see page 3)
 2. Morphine sulfate U.S.P. 15-20 mg ($\frac{1}{4}$ - $\frac{1}{3}$ gr) subcut or if necessary I.V. may be employed for the relief of pain. Perhaps meprobamate (Demerol®) 100-150 mg might be of value as a substitute for morphine sulfate because of its all-gentle antispasmodic action.
 3. Atropine Sulfate U.S.P. 0.4-0.6 mg ($\frac{1}{150}$ - $\frac{1}{100}$ gr) subcut should be given as an antispasmodic.
 4. Glyceryl Trinitrate U.S.P. (nitroglycerine) 0.3-0.6 mg ($\frac{1}{200}$ - $\frac{1}{100}$ gr) sublingually may be employed for relief of severe pain.
 5. Parenteral fluids
 - a. Plasma. Give 250-500 cc of plasma I.V. immediately and follow with subsequent infusion of plasma substitute as necessary to correct distended fluid balance.
 - b. Crystalloids. 5% glucose and/or normal saline may be used initially in lieu of plasma (when the latter is not available) to correct altered fluid and mineral imbalance.
 6. Withhold food and fluid by mouth.
 7. Employ continuous gastric suction.
 8. Careful observation. The patient should be constantly attended and vital signs should be checked at 15-30 minute intervals. As indicated during the acute period, blood count, hematocrit, serum amylase and lipase should be checked.
- B. Follow-up** After the patient has recovered from shock or if patient has not developed shock 2 alterations should be considered with regard to future immediate management.
1. Conservative or expectant management. This is to be preferred when verifiable. The patient should be observed closely for evidence of continuing inflammation of the pancreas and/or related structures. The opinion of a surgical consultant should be obtained in every case of suspected acute pancreatitis. Immediate surgical intervention. When the diagnosis is in doubt and there is a possibility of a serious and surgically correctible lesion (e.g., perforated peptic ulcer) an exploratory operation may be indicated.
 3. Observation. The course of the inflammatory process should be observed by frequent repeated physical examinations and blood counts and by blood sugar levels and serum and urine enzyme determinations as indicated.
 4. Supportive therapy
 - a. No fluid or foods should be given by mouth for the first 48 hours and continuous gastric suction should be maintained for that period.
 - b. After 48-72 hours small quantities of bland low fat liquid foods may be introduced gradually by mouth. If required gastric suction may be temporary. continued

several times during the day for small oral feedings and then gradually discontinued depending upon individual progress

c Fluid and electrolyte balance is maintained by appropriate parenteral fluid (page 7)

d Atropine sulfate 0.4-0.6 mg ($\frac{1}{150}$ - $\frac{1}{100}$ gr) s bcut may be administered t i d in an attempt to suppress pancreatic secretion

C C v l e s i C e W h i n t h e of pancreatic inflammation has cleared

1 Bland low fat diet should be given

2 Drug

a Belladonna extract 15 mg ($\frac{1}{4}$ g) t i d or atropine sulfate 0.4-0.6 mg ($\frac{1}{150}$ - $\frac{1}{100}$ g) t i d

b Antacid may be of aid (see page 254)

3 E l u s t f p a t e n t f o r g y C n s i d e r t h p a t i c fully relieved surgically treatment of biliary tract disease through proper treatment of fatty Sphincterotomy may be indicated in urgent patients of recurrent pancreatic inflammation

P o p h y l a x i s

A All associated logical factors should be considered e.g. biliary tract disease duodenal ulcer etc

B D i t P t i n t s w h o h a d p r e v i s m i d t t r k s o f u t p a e t i s h o u d b e p l d n l o w f a t d i e t a r y e t i m e a n d p r e m i t t d n o t h i s m a y r e d u c e t h e p o s s i b i l i t y f s u b e q u a n t a t t a k

CHRONIC PANCREATITIS (code No 690 956)

Chronic inflammation of the pancreas associated with fibrosis of the gland. In the interlobular type the external ducts are often dilated and digestive dysfunction. In the interacinar type the internal ducts are dilated and diabetes develops. A pancreatic carcinoma may develop. Pain at epigastric and periumbilical regions. Peptilorrhea, pathological digestion and general debility. Atherosclerosis may be associated. Urinary stones of pigment and fat. Flatulence and bowel irregularity. The physical examination reveals only an epigastric intermittent tenderness.

Labatory findings may include bulky foul fatty stool, indigestion of food, glycosuria, decrease of pancreatic enzymes and duodenal drainage. Pancreatic alkaline function may be absent.

T r e m e n t

A S p e c i f M N n
B G l M R m o a g g a t g f c t s w h p h i l

h p t b l y d g t r o d o d i d i s a s { g
p t a t g p p t i l) and al hol

2 Nutrit

Diet: High CHO low fat low protein high calcium
Diet: When pancreatic achylia is the conspicuous feature of the illness protein hydrolyses may be employed

supplement natural proteins. If diabetic present dietary modification may be necessary (see page 57).

- b Vitamins. Multivitamin tablets and B complex vitamins should be given.
- c Calcium salts. Calcium gluconate 1 Gm (15 gr) tablet 2-3 tablets tid may be given to help replace calcium lost in stool.
- d Replacement of deficiencies: pancreatic enzymes. Pankreatin (tablets) 0.32 Gm (5 gr) q valent to 15 g Pankreatin NF. Enteric coated 1-3 or more tablets tid per prn.
- e Detergent agent (e.g., solectan monocol) are of doubtful value in correcting impaired fat and calcium absorption.

3. Drugs

- a Oxbile Extract NF (bile salts) 0.5 Gm (7½ gr) tid per day be of value.
- b Diluted Hydrochloric Acid NF. B.P. 1040 (16-64 min) tid with meals.
- c Ferric Sulfate USP 0.2-0.3 Gm (3-4½ gr) tid per day for anemia.
- d Insulin for diabetes when present (see page 395).

PANCREATIC CARCINOMA (code No. 690.8)

Carcinoma of the pancreas occurs most commonly in males over 50 years of age. It is characterized by epigastric pain extending to the back rapidly and marked weight loss and multiple gastrointestinal complaints. Physical examination may reveal an epigastric mass, icterus and hepatic enlargement. Laboratory findings include evidence of disturbances of carbohydrate metabolism, elevation of serum lipase and amylase and widening of duodenal loop revealing S configuration of duodenum on x-ray.

Treatment

A. Nonoperative Measures. Symptomatic and palliative.

B. Surgical Measures.

- 1 Radical surgical resection in selected cases.
- 2 Palliative surgical operations. Biliary tract shunting procedure in individuals associated with jaundice.

Chapter 11

DISEASES OF THE URINARY SYSTEM

NONSPECIFIC URINARY SYMPTOMS

Urinary symptoms should never be ignored. Symptomatic treatment must never be substituted for a thorough investigation of the underlying organic or functional abnormality.

FREQUENCY OF URINATION (code No 706) (Nocturnal code No 707)

Frequency is one of the most common of the urinary symptoms and may occur either during the day or night. It may be caused by any of a variety of organic or functional disorders and is often of psychogenic origin.

If the symptom is disturbing to the patient, treatment can be instituted while diagnostic procedures are being completed. Use antispasmodic sedative drugs as for dysuria (see below). Fluid restriction may be employed particularly at night if there are no contraindications.

DYSURIA (code No 704)

Dysuria may be caused by infection of the genitourinary system or by lesions of the lower urinary tract. It is usually associated with urgency and frequency. Mild discomfort may also be produced by a highly concentrated acid urine.

Treatment

- A Specific Measures Treat the underlying disease
- B Symptomatic Measures Antispasmodic and sedative drugs
- 1 Atropine Sulfate U.S.P. B.P. 0.4-0.6 mg ($\frac{1}{150}$ $\frac{1}{100}$ gr) e. ry 3-4 hours or other parasympatholytic drugs (see page 34)
 - 2 Phenobarbital U.S.P. Phenobarbital B.P. 15-30 mg ($\frac{1}{4}$ $\frac{1}{2}$ gr) t. i. d. q. i. d. or more as needed
 - 3 Bladder sedative mixture
R Potassium citrate 30 0 3i
Hyoscyamus tincture 30 0 1
Elixir of phenobarbital q. s. ad 120 0 3iv
Sig. 4cc (1 dr) t. i. d. a. c. and b. s. or q. 4 H

OLIGURIA (code No 702) and ANURIA (code No 703)
(also see Lower Nephron Nephrosis page 303)

Oliguria and anuria usually serious symptom and may be due to many causes such as congestive failure dehydration renal failure or other less common disorders. It is important to differentiate these symptoms from urinary retention.

Treatment

A Specific Measures Treatment underlying disease

B Fluids Do not give excess fluids to patients with oliguria or anuria due to renal failure death will result from overhydration. Simplified hydration due to inadequate fluid intake in disorders (caused by severe peripheral diarrhea) usually corrected by oral or parenteral fluids. Replace any electrolyte (see page 15)

RETENTION OF URINE (code No 705)

Urinary retention due to obstruction may be the cause of hemorrhagic and most commonly distention of the prostate. The bladder may be percussed and palpated above the symphysis.

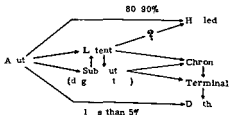
Place patient on micturition until the symptoms subside and attend by other means if necessary. If this is unsuccessful cannot be emptied the patient should be catheterized. Correction of underlying abnormality necessary to prevent recurrence.

SPECIFIC DISEASES OF THE KIDNEY

GLOMERULONEPHRITIS

Glomerulonephritis apparently different varieties predominantly involving both kidneys. The disease is believed by some to be a delayed allergic reaction but infection (usually bacterial hemolytic streptococcal infection) the bacteria and hemorrhagic form of the disease represent progression of the undischarged acute renal disease.

Though the disease is antibody mediated by the following chart



294 Glomerulonephritis

It is difficult in the initial attack to predict the course of the disease. However, about 80% of the patients have mild damage which allows for healing. Most of the other 20% enter the so called latent phase and the nature of the progression of the disease seems to depend primarily upon the extent of the renal lesion. The greater the amount of initial damage, the more rapid the progression to subacute, terminal stages and death. By proper treatment it may be possible to retard this process.

Diagnosis

Diagnosis of glomerulonephritis rests primarily upon the urinary findings of red blood cells and/or red blood cell casts; therefore, a careful examination of a freshly voided urine specimen is the best single examination in making the diagnosis.

General Principles of Treatment

The problems of therapy in renal disease are threefold. Treatment of each aspect of the disease will be discussed in terms of these principles:

A Addis's Principle of Rest. The cause of the progression of the renal lesion in glomerulonephritis is unknown. Addis suggested that progression is due to too great a work load for the amount of functioning renal tissue remaining. Most of the work of the kidney was assumed to be involved in the concentration of solutes (i.e., the reabsorption of water against osmotic pressure). Urea is the most important solute. Hence it was suggested that the loss of urea (from protein catabolism) and therefore the less work.

However, it has recently been shown that the percentage of energy utilized by the kidney (O_2 consumed) in the concentration of solutes is such a small fraction of the total as to make this thesis untenable.

On the other hand, the empirical evidence of Addis's principles has not yet been refuted. Therefore, an adequate but minimal protein intake remains important in influencing the course of the disease.

B Correction of Physiological Abnormalities. Since many of the manifestations of renal disease are associated with marked physiological abnormalities (e.g., hypoalbuminemia in the nephrotic syndrome), the therapy is aimed at correcting these physiological disturbances as they occur. However, many of the defects apparently corrected quickly revert to the disease state as soon as therapy is stopped (e.g., return of hypoalbuminemia after intravenous albumin is stopped). This may make continued intensive therapy imperative if one wishes to prolong life, especially in terminal cases. However, do not use therapeutic means which appear to correct the physiological defect but which are in themselves unphysiological or tend to defeat Principle A above (e.g., high protein diet to correct hypoproteinemia).

C Complications. Complications are treated as they arise and are discussed in the appropriate section.

ACUTE GLOMERULONEPHRITIS (code No 712 100)

235

Diagnosis

The history usually recalls an onset of hematuria, puffiness about the eyes, headache and occasionally in large and severe cases usually has a low specific gravity except when oliguria is present in which case it may be elevated. The urine sediment is loaded with red cells and white cells (tubular cells) but has only a few casts. Red cell casts are pathognomonic. Mild to moderate proteinuria is present. There may be increased blood volume and apparent or real anemia. The serum blood urea nitrogen and creatinine may be elevated. Hypertension is usually present.

Treatment

- A. Specific Measures There is no specific treatment. Treatment of acute nephritis with corticotropin (ACTH) or the cortisone has been generally quite unsatisfactory. A few cases respond well but these are exceptions. Corticotropin (ACTH) or the cortisones in this disease are of no value. Sodium restriction should be case of overdosage and added to increase edema and precipitate rigidly observed (see page 53). There is little that can be done to alter the abnormal physiology in increased blood volume elevated NPN etc. These tend to disappear spontaneously as the condition improves.
1. Home care & hospitalization The average case of acute nephritis can be cared for in the home although hospitalization becomes necessary if complications develop.
2. Bed rest The patient should remain in bed but may have bathroom privileges. The duration of the period of bed rest is difficult to anticipate but it probably is at least 2 weeks. A good rule is to keep the patient in bed until the sedimentation rate is normal until the blood pressure and blood NPN have been normal for at least 2 weeks or until the urinary findings show not more than 0.2 Gm protein/24 hour. Do not insist on bed rest if they remain for 11/2 months. Return to activity should be gradual.
3. Diet
 - a. I (1) Rye & light diet with protein restriction is indicated. Give 0.2 to 0.3 Gm protein/Kg (0.1 to 0.15 Gm/lb) body weight for first 7 to 10 days. Then increase to 0.5 Gm/Kg (0.25 Gm/lb) body weight plus the amount of protein lost in urine.
 - b. Fluids and potassium restriction. (5 to 7 Gm/lb) Adjust fluid intake to hanging physiological state (see page 53). If oliguria develops fluid restriction may be necessary to avoid drowning the patient. In the late phase of disease cannot always be induced by forcing fluids.
 - (3) IV fluids are rarely indicated except in cases a

associated with dehydration or abnormal fluid losses. In these cases it is important to give parenteral serum albumin 0.2 Gm/Kg (0.1 Gm/lb) of body weight and sufficient glucose to maintain closely to an isocaloric intake as possible. Give I.V. fluids slowly.

4. Blood. If anemia is marked, a carefully cross matched blood transfusion may be administered slowly.

C. Treatment of Complications

1. Cerebral edema with resultant headache and convulsions.
 - a. If headaches are not severe and convulsions are not present, give sodium pentobarbital 15-60 mg ($\frac{1}{4}$ to 1 gr) or paraldehyde 4-8 cc (1-2 dr) b.i.d. to q.i.d. as necessary.
 - b. If headaches are severe and convulsions are present, give magnesium sulfate 1.0 Gm (15 gr) (10 cc of 10% solution) I.V. slowly. *CAUTION: Whenever administering magnesium sulfate I.V. always have syringes filled with 10 cc of 10% solution of calcium gluconate or other calcium salt ready to administer I.V. if narcosis develops or respiration ceases.*
2. Cardiac failure. One of the most common causes of death in acute nephritis is acute cardiac failure. At the first signs of failure, digitalization should be instituted by one of the rapid methods (see page 197). Sodium restriction should also be instituted immediately.
3. Focal infections. Any focal infection, especially of an acute nature, should be treated promptly. Chemotherapy agents and antibiotics may be used as necessary for this purpose, but they are of no value in the therapy of the nephritis itself. Effective blood levels of sulfonamides and penicillin may be maintained with smaller dosages than under ordinary clinical circumstances; this is due to the decreased tubular excretion (especially of penicillin) of the decreased glomerular filtration (as with sulfonamides).

SUBACUTE GLOMERULONEPHRITIS (code No. 712.100.0) (Nephrotic Syndrome code No. 713.x40 or Degenerative Nephritis)

Dagnosis

A history of previous acute nephritis may not be elicited. The question of whether all cases of the nephrotic syndrome are preceded by acute nephritis has not been settled, but it appears that they are not. The most common physical finding is marked pitting edema. Proteinuria is marked, the urinary sediment may contain many atypical (especially fatty) casts, and many epithelial cells, a few red cells. Blood N.P.N. decreases slightly or may be low. The serum albumin is low and there is a marked lipemia. Anemia may also be found. Azotemia is also present sometimes.

Treatment

A. Specific Measures. Corticotropin (ACTH) and the corticosteroids have been employed in the treatment of subacute nephritis with

a. **Effect of the hormone** The use of these hormone results in a marked decrease or even disappearance of the albuminuria with a subsequent or concomitant diuresis and a gradual return of serum albumin to normal level. The dosage and duration of administration are quite variable. There are two main points of view.

1. **Intermittent therapy** The drugs are given in short courses of 7-14 days and repeated as necessary. Over 50% of patients treated have relapsed promptly (see page 423 for details).
2. **Continuous therapy** The advocates of continuous therapy (ACTH) and drugs insist that there is no point out that this is a chronic disease and that there is no evidence that these drugs in any way influence the ultimate course. The drugs are given continuously in the lowest dosage possible to keep the disease under control (e.g., urine free of protein).
All patients for administration of corticotropin (ACTH) and the corticosteroids should be observed.

B. General Measures

1. **Rest** The patient's rest for the kidneys is continued in this stage (see page 294). This stage may represent progression of the nephritis and leads into the terminal stage. The transition may be rapid or the recovery may be a matter of many years.
2. **Diet** Because of the massive albuminuria an attempt must be made to keep the body in nitrogen balance, but at the same time not to overload the kidney. There is no evidence that the hypoproteinemia can be corrected by a high protein diet per se. During long-term administration of corticotropin (ACTH) or the corticosteroids, great care should be taken that the protein intake is adequate—at least 2 Gm./kg. (0.5 Gm./lb.) per day.
Adults: 0.5 Gm. protein/kg. (0.25 Gm./lb.) body weight per 24 hours plus an amount of protein per 24 hours that is equal to the ill at the time. (Example: 70 Kg. man with proteinuria of 10 Gm./day = 0.5 Gm./kg. body weight = 35 Gm. protein = 10 Gm. lost in urine = 10 Gm. protein. Total = 45 Gm. protein per day as the approximate intake.)
b. Adults: 0.75 Gm. protein/kg. (0.35 Gm./lb.) body weight per 24 hours plus urine loss.
3. **Treatment of edema** The physician of the physician's physical judgment of view that is accepted as a guide. For this the renal disease is a liability.
Sodium restriction. The probably the method of this is through oral therapy. It is when used alone. The sodium restriction is 10 Gm. (15 gr.) per day. It is a restriction to 0.5 Gm. (7½ gr.) per day may be necessary. This restriction should not be carried longer than is clinically necessary and when the restriction is marked the patient should be watched for symptoms of sodium deficiency. Salt substitutes should not be employed. Educate patient to live on low salt intake. Long term use of exchange resins has proved an satisfactory, they are unpalatable and tend to produce indigestion.

b Mechanical removal Whenever the fluid accumulation becomes very marked mechanical removal is one of the most beneficial methods This includes removal from pleural and peritoneal cavity and especially the use of Southey's tubes to remove massive edema from the legs Any infection resulting from use of Southey's tubes should be controlled with antibiotic agents

c Agents to increase osmotic pressure of blood

(1) Salt poor human albumin Of all the measures that have been employed to increase osmotic pressure this agent has the soundest physiological justification 50 Gm (1 2/3 oz) per day I V may induce a rapid diuresis However the effect is transient most of the albumin is lost in the urine and much of the remainder is rapidly catabolized After cessation of the therapy (may be continued for from several days or weeks) there is little evidence that the course of the disease has been modified and in most cases the serum albumin concentration will return to its former low level

(2) Blood plasma Plasma has little value primarily because of its high salt content It may also carry the virus of infectious hepatitis

(3) Some of the newer plasma expanders (Dextran® Gelatin Plazmoid® etc) have been employed Although they may induce a temporary diuresis their routine use is not indicated

(4) Other preparations such as acacia and isinglass are mentioned merely to be condemned Their use is entirely unphysiological

4 Chlorothiazide (Diu il®) in dose of 0.25-1 Gm (4-15 gr) daily appears to be a promising diuretic agent in this condition

5 Acetazolamide (Diammonium chloride) These drugs may be used but their effect is often not noted until mild acidosis develops Since these patients may readily develop acidosis caution should be exercised in use of these drugs

6 Mercurial diuretics are not advised They may cause at least temporary renal damage and generally are not beneficial

7 Water Patient should be encouraged to drink adequate fluids As long as there is sodium restriction water will not accumulate in the tissues Forcing fluids however is of little value in inducing a greater diuresis if fluid intake and urinary output are adequate

8 Induction of infection It has long been known that patients with the nephrotic syndrome develop infections following some virus infections especially measles In susceptible children with subacute nephritis exposure to one of the mild exanthematous diseases may be indicated

C Treatment of Complications The principal complications are infections commonly pneumonia and pneumococcal peritonitis These should be treated with the appropriate chemotherapeutic and antibiotic drugs

LATENT GLOMERULONEPHRITIS (code No 712 190)

The patient with latent nephritis may or may not give a history of an attack of acute glomerulonephritis. In the latent phase although there are no complaints or physical findings the lesions either have not healed at all or there is insufficient healed tissue to carry the entire load of work. *The latent phase may last for as long as 20 to 30 years* and the patient may die of other intercurrent disease before his renal function fails.

Diagnosis

The only positive findings are occasional red blood cells and cell casts and transient to persistent albuminuria. The physical examination and all blood findings (hematological and chemical) are normal.

Treatment

A General Measures

- 1 Diet The patient should be on a minimal but adequate diet containing 0.5-0.75 Gm protein/Kg (0.23-0.35 Gm/lb) body weight. At least 50% of the protein should consist of dairy products, vegetables and cereals.
- 2 Fluids The patient should be taught to follow fluids 3,000-4,000 cc (3-4 qt) per day.
- 3 Activity The patient should be cautioned against strenuous exercise but should be encouraged to live as normal a life as possible.
- 4 Physiological Consideration Since there is no apparent physiological abnormality no specific treatment is indicated.

- B Complication** Psuedo-exacerbation of symptoms. Patients with latent nephritis have a characteristic nonspecific response to any febrile illness; this is particularly marked in the case of foreign protein reactions (e.g. vaccination, inoculation, or infections). This reaction is characterized by hematuria (often gross) and a mild increase in proteinuria and white blood cell coming on immediately with the fever and subsiding with the fever. *This is not another attack of acute nephritis.* There is no delay between infection and renal reaction; no hypertension, no edema, and no anuria.

It is exceedingly rare to have a second attack of true acute glomerulonephritis. Most cases of so-called second attacks are really exacerbations of latent nephritis. These exacerbations never damage the kidneys as severely as the initial attack and one can rarely detect any change in renal status after the attack is over.

- 1 Prophylaxis Because of the association of exacerbation with fever, infection and vaccination, one should avoid these insults whenever possible. Patients with latent nephritis should not undergo vaccination routinely.
- 2 Treatment There is no treatment of the renal lesion other than continued treatment of the latent nephritis. Treatment is indicated entirely at the precipitating cause. The patient should be kept in bed for about 1 week after fever has disappeared and should be allowed up slowly over the next week.

CHRONIC OR TERMINAL GLOMERULONEPHRITIS (code No 712 100 0)

It is difficult to say when the terminal or chronic stage begins. It is the time at which signs and symptoms of renal insufficiency develop. It may be very difficult to detect early, but as it develops certain findings appear. Most characteristic are the (1) elevation of blood N P N (2) development of anemia (3) gradual elevation of blood pressure and (4) presence of a few casts and red blood cells in the urine. However, the blood protein is normal, edema is absent early and there is slight proteinuria. This stage may last from several months to a few years.

Diagnosis

A history of acute or subacute nephritis may be elicited. The physical findings vary with the severity of the disease, but hypertension with its associated vascular changes is the most common finding. Edema usually appears and may be due to cardiac or renal failure. The urine has a low or fixed specific gravity. There is a mild to moderate proteinuria. The sediment contains a few red cells and broad casts (renal failure casts) and a few epithelial cells. As anemia develops, increased blood urea nitrogen, alterations in electrolyte balance and a decrease in serum proteins occur.

General Treatment

- A Diet. As the blood N P N and creatinine begin to rise, any increase in protein intake is followed by a marked rise in N P N. The patient's protein intake must be restricted to 0.5 Gm /Kg (0.23 Gm /lb) body weight, plus the urinary losses.
- B Fluids are forced to 3,000-4,000 cc (3-4 qt) per day.
- C Treatment of Physiological Abnormalities. As the N P N continues to rise and renal failure becomes more severe, there is a progressive tendency to acidosis and altered electrolyte balance. The kidneys become unable to form ammonia or conserve fixed base and fixed base elements consequently begin to decrease in the blood.
 - 1 Early in the terminal phase these are replaced by oral use of salts. Either of the following may be used:
 - a Calcium lactate or chloride 3.5 Gm (45-75 gr) daily
 - b A mixture of the following salts:

Rx Sodium	1 tr	100.0	3xxv
Calcium	chlorid	3.0	g vi

 Sg 2 Gm (30 gr or 1st tsp) in 1 glass water t.i.d.
 - 2 Alkalizing urine. The urine should be maintained at a pH greater than 6.0 with sodium citrate or sodium bicarbonate 1-2 Gm (15-30 gr) q.i.d. This is done to help prevent cast formation in the collecting tubule.
 - 3 Hospital treatment. As uremia becomes more marked and acidosis more profound, nausea and vomiting develop, it is generally necessary to place the patient in the hospital in order that the electrolyte balance may be adjusted as needed with I.V. fluids (see page 19). Uremia must also be treated (see next page).

HEALED GLOMERULONEPHRITIS

Any patient who has had an attack of acute glomerulonephritis has undoubtedly suffered permanent destruction of some of the nephrons. The lesion is said to be healed when there is no longer any evidence of activity and the number of remaining functioning nephrons is great enough so that no impairment in function or structure can be found. However, there is no way of estimating how many nephrons this may be. It is always possible that the number functioning is barely sufficient to satisfy the average demands of the body.

Follow up Care

Subsequent diseases may cause sufficient additional nephron damage to bring about a latent glomerulonephritis. Patients with a healed glomerulonephritis must therefore submit to a moderate but not necessarily rigorous protein restriction and should have urine examinations at least once a year for life.

UREMIA (code No 551)

Uremia is a physiological state resulting from renal insufficiency which may be defined as an alteration in electrolyte balance with retention of nitrogenous and other waste products. Although uremia is most frequently seen in the terminal phase of chronic renal disease it does not necessarily imply an early demise. Some cases of uremia may actually be cured, e.g. those resulting from urinary retention secondary to obstruction. In the management of uremia one should remember that the alterations in electrolyte balance are more important than the elevation of the B.P.N. and that therapy therefore should be aimed primarily at preventing and treating the acidosis which develops.

Pathological Physiology of Renal Insufficiency

A. Renal Defect

1. Glomerular filtration is depressed and produces an elevation of the serum N.P.N. phosphorus, sulfate and other acids. Also this leads to metabolic acidosis (see page 18). The serum phosphate falls and the serum calcium tends to fall.

2. Tubular function is depressed and the kidney loses its power to manufacture NH_4^+ (which combines fixed bases); this leads to loss of the fixed bases and urea, potassium and calcium which further contribute to acidosis (see page 18).

B. General Metabolic Effect. Anemia develops gradually due principally to bone marrow depression. Loss of calcium in urine with consequent low serum calcium and high serum phosphate leads to parathyroid hyperplasia. Parathyroid hormone can not be secreted however and remains elevated. The serum phosphate may become low or d.

C. Gross

The altered physiology leads to the clinical manifestations of uremia. These are variable. In early uremia there may be lethargy

headache pruritus and weakness. Late uremia is characterized by acidosis and dehydration. In addition tetany may result from lowered serum calcium and muscular weakness may occur if serum potassium is lowered. The blood N P N sulfate and phosphorus are elevated the serum potassium is variable and the serum sodium calcium and CO_2 are lowered. A normocytic anemia is present. Coma is superimposed later.

Treatment

A Early

- 1 Diet Protein must be restricted to 0.5 Gm./Kg. (0.23 Gm./lb.) body weight plus the amount lost in urine. This tends to reduce N P N and serum sulfate (see page 294).
- 2 Fluids and electrolytes Force fluids orally to 3,000-4,000 cc (3-4 qt.) per day. Give calcium lactate and salt mixture by mouth as for terminal glomerulonephritis (see page 300). This helps keep the electrolytes in balance.

B Late

1 General measures

- a Diet As above. Protein restriction is very important.
- b Fluids
 - (1) Force fluids orally to 3,000-4,000 cc (3-4 qt.) daily unless patient is anuric.
 - (2) I.V. fluids and salts should be given as necessary to maintain normal electrolyte balance (see page 15).
- c Electrolytes
 - (1) Continue use of salt mixture (see page 300).
 - (2) Aluminum hydroxide gel 15 cc (4 dr. or 1 Tbsp.) q.i.d. orally aids in reducing the hyperphosphatemia (causes precipitation of insoluble phosphates in bowel) and so helps to elevate serum calcium and prevent tetany.
 - (3) Calcium gluconate or lactate 10% 10 cc (2½ dr.) I.V. is useful primarily to treat or prevent tetany.
- d Transfusions of carefully matched whole blood or red cells may be used to control anemia. All other forms of treatment to combat anemia are without benefit.

- 2 Complications of treatment In the treatment of uremia the physician is apt to encounter a therapeutic dilemma. In the course of attempting to correct the electrolyte balance the amount of sodium that must be administered may cause the patient to develop cardiac failure. Little can be done for the patient at this time as he has almost no cardiac or renal reserve remaining.

C Terminal

- 1 Calcium lactate or gluconate 10 cc (2½ dr.) of 10% solution I.V. primarily to control tetany and convulsions.
- 2 Magnesium sulfate 1 Gm. (15 gr.) (10 cc of a 10% solution) I.V. primarily for reflex spasms and convulsions. Caution: Have I.V. calcium salts ready in syringe (see page 295).
- 3 Paraldehyde 20 cc (5 dr.) in 30 cc (1 oz.) of oil rectally or 4-8 cc (1-2 dr.) I.M. as necessary for sedation.

EXTRARENAL AZOTEMIA

Extrarenal azotemia is the abnormal accumulation of nitrogen waste products in the presence of normal or potentially normal renal function. The most common cause is a decreased effective circulating blood volume with inadequate glomerular filtration as happens in shock and dehydration, etc. It also occurs in massive gastrointestinal bleeding, whether it is a sudden excessive protein digestion and absorption, decreased circulating blood volume.

Treatment

Treatment is aimed entirely at correcting the underlying condition, not renal disease. If present, fluids and electrolytes suffice to restore the blood chemistry to normal should be given.

ACUTE RENAL FAILURE

(Lower Nephron Nephrosis code No 713 y00 9)

(Due to Hemoglobinemia Following Transfusion code No 713 38x 9)

Pathological Physiology

It has been demonstrated that the cause of renal failure (oliguria, anuria) which occurs in a variety of toxic conditions present in the same clinical and pathological picture is a variety of the etiology. This condition is most often induced by one of the following: (1) intravascular hemolytic reactions (e.g., transfusion reactions), (2) chemical injuries, (3) burns, (4) chemical toxicity of some types (e.g., acute tetrahydro sulfonamides, etc.), (5) toxemia of pregnancy, and (6) non-traumatic renal ischemia. Although pathogenesis is variable, the histopathological picture is the same, primarily focal glomerular necrosis of the distal convoluted tubule with blood stasis in the lower nephron and interstitial edema.

In mild to moderate cases the kidneys will open spontaneously in 1 to 14 days (if the patient can be kept alive that long). In more severe cases, renal shutdown may be permanent. The evidence suggests that if the patient survives, recovery is complete and that the possibility of the kidney may occur in as short a time as 2 to 4 weeks.

Diagnosis

- A Prodrome of Shock. At the onset, symptoms of shock may be the only finding. Hemoglobin may be found in urine.
- B Prodrome of Renal Shutdown (Myoglobinuria or Myoglobinuria). The patient may be symptomatic but without obvious peripheral edema. The rules of pulmonary edema may be found if the patient becomes overhydrated due to over-treatment with fluids. This drowning is the most common condition.
- C Prodrome of Recovery. The diuresis which follows renal shutdown may be marked and uncontrolled and may lead to dehydration. Muscles weaken (detest low potassium) and tetany (due to low serum calcium) may occur. The blood NPN usually does not return to normal until 2 to 4 weeks after initial recovery of the kidney has occurred.

TreatmentA Emergency

- 1 **SHOCK** Since many cases are associated with traumatic or burn injuries the renal ischemia associated with shock may play a role in the pathogenesis. Immediate and vigorous anti shock therapy is important (see page 27)
- 2 Immediate alkalization of the urine in cases of transfusion reaction may help prevent the precipitation of acid heme compounds in the renal tubules. Give sodium bicarbonate 5-10 Gm (75-150 gr) orally at once. Check the urine pH every 1-2 hours and give sufficient sodium bicarbonate to keep the urine alkaline.

B Oliguric or Anuric Phase Management in this phase is very difficult and should be undertaken only by trained personnel in a hospital able to determine chemically the entire electrolyte panel. (See page 15)

- 1 **Weight patient accurately daily.** Weight gain means fluid retention and this must be avoided. These patients should generally lose 0.3-0.5 Kg/day which represents endogenous tissue catabolic losses.
- 2 **Fluid restriction.** This is one of the foremost principles in the therapy. In the past patients were often drowned to death in an effort to promote diuresis. Usually 800-1500 cc of fluid is the maximum allowed daily. If patient is not losing excess fluids (as by vomiting or diarrhea or excess sweating with fever) the insensible water loss plus urinary loss is the only fluid which must be replaced. The insensible loss can be calculated as 15 cc water/Kg per day. However, some water is supplied from oxidation of food calories. This may average as much as 400-500 cc per day. Therefore the usual fluid requirements are 400-600 cc per day for a 70 Kg patient. This may be taken orally. If vomiting occurs the fluid may be given I.V. as 10-15% or more concentrated glucose given slowly and carefully to avoid a butanous effect at once. The patient should never be allowed to gain weight (keep an accurate record of weight) for this probably represents fluid retention. If the patient is vomiting has diarrhea or is sweating give additional fluids cautiously to replace the loss.
- 3 **Electrolytes.** In the absence of vomiting or other extrarenal losses no electrolyte replacement is needed. The electrolyte pattern should be examined daily and every attempt made to keep the electrolyte values within normal ranges. Give electrolytes as needed orally or parentally. In most cases potassium either in food or as electrolytes must be avoided. Give calcium gluconate 10% (2½ dr) 10% solution I.V. for convulsions.
- 4 **Diet.** A high carbohydrate and high caloric diet without protein will prevent endogenous protein breakdown and slow down the accumulation of protein breakdown products (i.e. urea, organic acids and potassium). In the absence of vomiting a simple way to supply fluid and food is as follows: Pass a small polyethylene plastic tube intrasally into the stomach. Calculate the amount of fluid necessary over 24 hours and to this add lactose and cod liver oil to give the number of calories required for maintenance. The mixture

may then be emulsified in a 1:1 d r by adding 2.5 of Tween 80® to the solution. The volume is added to 24 parts and a hypodermic syringe given hourly through the polyethylene tube. If the patient is vomiting 50% or less concentrate d glucose can be given by I.V. drip. Vitamins should be given with the high CHO intake either orally or I.V.

- 5 Digital ton If iden f a d a f i l o n l g ment
o p i n o n r y e d e m d e l o p m e n t e r a p i d d i g i t i z a t i o n
a d m a n t e s h o u l d b e c a r d o t (e e p g 197)
- 6 D e s s I f d r e s h o t o d b y t h 12th 14th day
i m a y b e n a y l o o t t m e d a t m e s s t o
o m b t t h e i s g a o t e m a d e s p i a l l y t h e h y p k a l m i a
A m o n g t h o e w h h a e b e a d o c a t e d t h t i f l k d n e y
b t O r a l e c h g a s i t e s t l i v a g a d o r
p a r t o n l i r r g a t i n a r e d i f f i c u l t t o u s b t m t a t t m e s
b e s d f i f a l k i d e y n t a y l i b l

C Reov y (D r t) Ph The managem t of th d t c
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m e t h d o t p a t l f t h p t e t N + o w t u n t k d r
g t h l g i p h s h b e e e s s I f t h r i t
b e s e d t h p t e t s h l d h v d a l y s e m N a + K + a n d C l
d t m a t i o s T h f l d a n d l t o l y t e r p l c m t
d t e m e d t h e b a f t h e s m u e m e t (e p g 15)
D e s s m v o n o o f t h r w y

- 1 Diuresis with isotolytic loss. The electrolyte disturbance is generally characterized by an excess of anions of sodium chloride and potassium in the urine with rapid fall of these ions in the cellular fluids. It may be necessary to give 20-30 Gm (2/3 l) of sodium chloride per 24 hours to maintain the balance. Urine is often death may result from hypochloremia and dehydration. Potassium chloride may also be given.
- 2 Diuresis with isotolytic retention. In this type is the appearance of an enormous retention of the enditubule in which sodium and chloride are absorbed in large amounts. In this condition is a hypernatremia. Na and Cl with excess of diuresis. The pyrexia is of immediate retention of Na^+ and Cl^- and the loss of these fluids.
- 3 Diuresis with osmotic isotolytic

Follow Up

When you complete the timeline review

INFECTIONS OF THE URINARY TRACT

Inf tions of the in y t act m y inv lv on or m e p ri
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organisms may also cause infection. Diagnosis is usually suggested by the presenting symptoms and signs and confirmed by microscopic and bacteriological examination of a sterile urine specimen including stained smears of sediment and colony counts of the urine. 25% of cases of infections may have bacilluria without pyuria.

A chronic or recurrent infection, particularly if resistant to antibacterial agents, suggests obstruction and urinary stasis. The final clearing of such infection is dependent upon the correction of the obstruction.

General Principles of Treatment

- A Correction of structural abnormalities which produce stasis is of utmost importance in cases with remediable defects. Urinary tract infections may disappear spontaneously or be easily cured as soon as the defect is corrected. The permanent eradication of infection in the presence of such obstruction is usually impossible. The diagnosis of obstruction usually requires cystoscopy and/or excretion or retrograde pyelography. Treatment is generally surgical.
- B Treatment of the infection with suitable chemotherapeutic or antibiotic agents as determined by bacteriological studies.
 - 1 Careful examination of fresh sterile urine specimen (2nd glass specimen in male catheterized specimen in female) for presence of pus and Gram's stain for preliminary identification of organism.
 - 2 Bacteriological identification of organism and determination of sensitivity of organism to antibiotic agents whenever possible. The latter is of special importance when streptomycin, chloramphenicol (Aureomycin®) or oxytetracycline (Terramycin®) are to be used because adequate dosage must be assured to eradicate infection before organism resistance develops (see page 514).
 - 3 Careful follow up to diagnose and prevent development of chronic infections.

INFECTIONS OF THE KIDNEY

Diagnosis

The manifestations of all infections of the kidney are similar but they vary in intensity with the severity of the infection. Symptoms include lumbar pain which usually radiates into the lower genitourinary tract but may radiate elsewhere. Chills, fever and nausea and vomiting as well as frequency, urgency and dysuria. There is usually moderate to marked costo-vertebral angle tenderness. Examination of a sterile urine specimen for pus and organisms is necessary to make the diagnosis and to select the proper antibacterial agent.

- A Pyelitis (code No. 722.100). Simple infection of the renal pelvis which does not affect kidney function.
- B Pyelonephritis (code No. 719.100). Renal infection which depresses kidney function and which in its chronic form may produce effects similar to those of chronic glomerulonephritis.
- C Pyonephrosis (code No. 722.100.2). Renal infection of greater severity than pyelonephritis with pus in the renal pelvis.
- D Renal and perirenal abscesses are surgical renal diseases.

TreatmentA Specific Measures

- 1 Antibacterial therapy should be given as soon as causative organism is identified and as soon as sensitivity tests have been conducted to determine dosage (see page 514)
- 2 Surgical treatment of any remediable obstruction should be carried out when the acute symptoms have subsided. Diagnostic studies of the urinary tract should be deferred also until the acute phase has passed.

B General Measures

- 1 Bed rest until completely asymptomatic
- 2 Fluids If the kidney function is not depressed and there are no other contraindications fluids should be forced. Minimum daily urine output of 1500 cc or more.
- 3 Analgesic and sedatives necessary for the comfort of the patient.

C Treatment of Chronic Pyelonephritis There are a series of chronic pyelonephritis in which the kidney has been moderately to markedly damaged. Infection in these kidneys is very difficult to eradicate. Additionally suggested the continuous use of small doses of sulfonamide drugs 100-200 mg (1½-3 gr) tid qid (after other measures have been taken to eradicate the infection) in the hope that the small doses might stop or slow down the progress of the disease. Once this therapy is begun it should probably be carried on for life.

Treatment of pyelonephritis is handled the same as terminal glomerulonephritis (see page 300).

CYSTITIS

(Acute code No 730 100) (Chronic code No 730 100 0)

Definition

Inflammation of the bladder is many times more common in women than in men and is more commonly due to Escherichia coli. It must be differentiated from urethritis which has similar manifestations.

A Symptoms Mainly dysuria, urgency and frequency. If severe there is tenderness and at times urinary retention. Chills and fever may occur. When infection is very severe hematuria may develop.

B Signs Suprapubic tenderness may be present.

C Laboratory Examination

- 1 Organisms and pus will be found in the urinary sediment if properly collected specimen.
- 2 Organism may be identified by examination of the smears (methyl blue-gram) and by culture for proper selection of the antibacterial agent (see page 514).
- 3 Two glass tests may be used to differentiate urethritis from cystitis in the male. Examine the urine grossly and microscopically. If the urine in the condensation is turbid infection is in the urethra. If all urine is turbid the bladder is infected (U-3 collection beakers or glass).
 - a First glass on lot of 4 cc of urine and contains the elements from the urethra.

- b Second glass contains the remainder of the urine from the bladder
- c A third glass may be collected after prostatic massage. In this method the patient must retain some urine in the bladder to wash out any residual material.
- D Cystoscopy May be necessary to determine the presence of obstruction, upper urinary infection, or source of bleeding. This must not be done during the acute phase.

TreatmentA Specific Measures

- 1 Antibacterial agents. Select the appropriate drug by bacteriological examination and sensitivity tests (see page 514).
- 2 Surgery. Correct any removable obstruction after the acute stage has subsided.

B General Measures

- 1 Bed rest if severe.
- 2 Fluids. If urination is painful, fluids should not be forced. When dysuria has subsided, maintain a high urine output.
- 3 Bladder sedatives and analgesics.
 - a For severe pain. Mild local anesthesia can be obtained by bladder instillation of 2% solution of Procaine Hydrochloride U.S.P. (Metycaine®) or 1:1000 (0.1%) solution of Dibucaine Hydrochloride U.S.P. (Nupercaine®). Allow the anesthesia to remain in the bladder for 10 minutes by placing a clamp on the catheter. After draining off the anesthetic, instill 10 cc of 5% solution of mild silver protein or 1:10,000 silver nitrate solution and leave in the bladder.
 - b For tenesmus. Treat as for dysuria (see page 292).

TUBERCULOSIS OF THE URINARY TRACT

(Kidney code No 710 123) (Bladder code No 730 123)

Chronic tuberculosis of the urinary tract usually occurs first in the kidney and involves the bladder secondarily. A history of bladder irritation is usually present; the urine contains pus and a few red cells, but there is generally no organisms. The urinary sediment must be examined microscopically and bacteriologically (culture and guinea pig inoculation) for acid fast bacilli. If tubercle bacilli are found, determine the primary urogenital site of the infection and whether renal disease is unilateral or bilateral.

Treatment

- A Treatment of renal tuberculosis with the newer anti-tuberculous chemotherapeutic agents tops prognosis and may effect cures in some cases. Therapy is the same as that advocated for pulmonary or other systemic tuberculosis. The use of intermittent streptomycin plus aminosalicylic acid (PAS) and/or isoniazid for 1 to 3 years has been advocated (see page 133).
- B Surgery. If unilateral tuberculosis is found and if the kidney is seriously involved, nephrectomy with subsequent streptomycin therapy should be considered. This would apply to help to

could be any low test tube cultures that may be present
C Symptomatic and supportive measures as necessary

OTHER DISORDERS OF THE URINARY TRACT

CARCINOMA OF PROSTATE (Adenocarcinoma code No 764 8091)

Diagnosis

The vast majority of prostatic inomas can be diagnosed by the finding of hard glandular areas than the gland on rectal examination. Confirmation of the diagnosis is made by needle biopsy and the finding of an elevated acid phosphatase. X-rays of the pelvis and lower spine are taken to determine the presence of metastases.

Treatment

A Early Cases Treatment of the early radical surgical removal. This is done only for those in which (1) there is no evidence of metastases (2) the gland is not enlarged (3) the patient is otherwise a good surgical risk and has a good life expectancy.

B All Other Cases

1 Hormonal therapy. The androgens have been found to be of great benefit in relieving the pain associated with metastases and in arresting the progress of the disease (in some cases totally causing a regression). The wide testosterone has been given with Dihydrotestosterone (USP) Stilbestrol (BP) and Ethinyl Estradiol (USP).

a Dosage

- (1) Androgen maintenance dose Dihydrotestosterone 1 mg ethinyl estradiol 0.1 mg per day orally if life.
- (2) If response is poor may increase to Dihydrotestosterone 2-3 mg ethinyl estradiol 0.2-0.3 mg per day.
- (3) Some of the effects of the progestins should be noted. Dihydrotestosterone 5-10 mg ethinyl estradiol 0.5-1.0 mg per day.

b Toxicity

- (1) Nausea and vomiting may occur but usually disappear when the drug is continued. If the cumulative dosage and increases tolerance develops.
- (2) Prostatic enlargement develops usually develops on continuation but this is not a indication of tumor development.

- 2 Orchiectomy. The result to date suggests that bilateral orchiectomy is the best method of great value in the treatment of prostate cancer.
- 3 General supportive measures such as general mobilization postoperative but attention to should be emphasized.
- 4 Bilateral orchiectomy and androgen therapy (with testosterone and estrogen maintenance) in order to improve quality of life must still be considered as a permanent treatment.

most cases seem little better than orchiectomy plus estrogens alone

UROLITHIASIS

(Renal Calculus code No 719 615) (Ureteral Calculus code No 723 615) (Renal Colic code No 711)

Renal colic is usually caused by the passage of a renal calculus into and down the ureter. There is a sudden onset of severe lumbago pain on the affected side radiating to the groin bladder testis inner thigh or other adjacent areas. The pain requires narcotics sometimes in large doses for relief. Nausea and vomiting may occur but no other constitutional symptoms represent unless there is a pre-existing infection. Urine output is reduced and hematuria is common. Stones may pass without symptoms.

Treatment

A. EMERGENCY MEASURES

1. Narcotic for relief of pain. This may have to be repeated if pain is severe.
 - a. Morphine sulfate or hydrochloride 15 mg ($\frac{1}{4}$ gr) I.V. or subcut. Stat. Atropin sulfate 0.5-0.75 mg ($\frac{1}{80}$ - $\frac{1}{60}$ gr) may be given with the morphine.
 - b. Meprobamate Hydrochloride Injection U.S.P. (Demerol® Dolantin®) 0.100 Gm ($1\frac{1}{2}$ gr) I.M. or orally in place of morphine. This has a minor atropine like effect in addition to its narcotic action.
2. Heat over the affected flank and lateral abdominal area may give some relief. This can be given as warm moist towels heat pad or warm tub bath.

B. General Management

1. Fluids. If patient does not develop anuria or oliguria fluids should be forced in order to maintain a high urine flow. Give fluid I.V. if vomiting prevents oral administration.
2. Check carefully for passage of stone. If this does not occur examine by x-ray for position of stone.
3. Attempt to remove stone by having patient void through a funnel layered with several thicknesses of gauze and analyze chemically to determine type of stone (calcium phosphate uric acid cystine etc.).

C. Surgery. If a stone becomes lodged in the ureter it should be removed surgically to prevent hydronephrosis.

D. Coexisting infection should be treated with suitable antibiotics (see page 514).

Prophylaxis

A. Correction of Underlying Disease. Treat any disorder which may cause or contribute to formation. These include hyperparathyroidism (see page 377) urolithiasis urinary infection (see page 306) gout and rarely cystinuria. Every patient with urinary tract calculi should have at least one serum calcium and phosphorus determination to rule out hyperparathyroidism.

B. Fluid. Any patient who has had a urolithiasis should drink large amounts of fluid at all times. In cases of urolithiasis the urine should be kept alkaline if possible.

Chapter 12

DISEASES OF THE MUSCULOSKELETAL SYSTEM

INTRODUCTION

Classification of Rheumatoid

- 1 Arthritis due to specific infection as tuberculosis
- 2 Arthritis due to humoral fever
- 3 Arthritis rheumatoid
- 4 Degenerative joint diseases
- 5 Arthritis due to direct trauma
- 6 Arthritis due to gout
- 7 Other thropathies (due to congenic neoplasms, metabolic, vascular, hematologic, non-allergic, toxic and unknown causes)
- 8 Fibrositis, myositis, bursitis

Examination of the Patient

The examination of the patient with rheumatoid arthritis should include a careful history and physical examination with special emphasis on determining the functional status of the joint (range of motion, ankylosis, deformity, atrophy, etc.). Routine laboratory studies include sedimentation rate and x-rays of the most involved joints. Initial to complete the diagnostic picture. Additional studies may include determination of the blood uric acid, sedimentation and examination of joint fluid and cultural immunologic and other tests for specific infections. The studies are important not only for making a differential diagnosis but also for providing a baseline for planning the therapy and evaluating the clinical progress of the patient.

The differential diagnosis of the four forms of arthritis is to be found in the table on page 312 and 313.

RHEUMATOID ARTHRITIS (code No. 24.1x0)

Rheumatoid arthritis (a profoundly humeral disease) is a chronic debilitating disease of undetermined origin. It is ordinarily a slow developing primarily the joints but it is actually a process of involving most of the soft parts of the body particularly the connective tissue, malodorous in the lymph nodes, bone marrow, spleen, gastrointestinal tract, endocrine system, eyes, parathyroid glands, connective tissues and the musculoskeletal system. The disease may involve any or all joints and is of varying severity. Peritendinitis is prominent in the

DIAGNOSTIC CHARACTERISTICS OF THE MAJOR FEATURES OF ARTHRITIS

	Rheumatoid Arthritis	Arthritis Due to Specific Infection	D degenerative Arthritis	Arthritis Due to Gout
Family history of similar condition	+		+	+
Past history	Frequent infections	History of specific infection		
Sex	Most common in women	Either sex	Both sexes	Usually men
Age at onset	Any age usually 20-50	Any age	Usually over 40 years	Usually over 35 years
General physical condition at onset	Poor undernourished	Acute good	Good but may show other senile changes	Good
Type of onset	Insidious (subacute)	Chronic may be poor	Insidious (slow)	Sudden (cessation of symptoms also sudden)
Feeling	+	Acute infection sudden		+
Joints involved	Any joint if symmetrical with tendency to spread centripetally Proximal finger joints especially involved	Chronic infection slow + (especially acute)		(during acute episodes)
Periarticular swelling	+	Any joint pyogenic forms are usually monarticular Non pyogenic forms are often polyarticular	Usually the large and weight bearing joints Also distal joints of fingers	Any joint monarticular or polyarticular Especially involves metatarsophalangeal joint of great toe
Ankylosis	+	+		+
Muscle atrophy	+	+ (pyogenic)		
Deformities	+	+ (pyogenic)		+ (late)

Cutaneous changes	Skin over joints and glomy	Similar to rheumatoid arthritis	Seile changes	Ely N n Late Similar t heuma told a thritis.
Sbcut od le	+			+ (tophi)
Ami	+	{ ly } + {chron}		
Luk yto i	±	±		± (du ing a te p od)
Blood s dime ta ti	+	+		+ (du g acut plisode and chr ni ph e)
Hype uricemia (blo d ic a id)				+
Char t r f joint fluid	Non pu lent (ste il)	Purul nt n purul t (s pti)	Non p ul nt (il)	N pu ul nt (t il)
X ray appearance f joints	Early Gn lized de cal ill tion of bones and joint effu i n Late Nar owing of joint space bon d st ti ankylosis	Sum i r to rh um t d arthritis but bony de calcifi tion more p minent near in volved joints	No hanges until late tag auses lipping osteophyt a and nar wing of joint p e	Early Normal Late Punched out appearan
Sp ific therapy	Peripheral G ld alite Spinal (spondylitis)	Sp if ant Inf ti agent	None (l y l t s ?)	C l h c in
Other ds g tic se tures		B t ological and immunological evi dence of pe ific local o systemic infe tion	Some lini lars in clude m n pausal arthritis in this category Estrogens are f value in e of gonadal d ficiency	1) E rly in di a e be tw n episode p tient is asymptomatic and ther is no resid l joint involvement 2) Tophi C ntain urate cry tal

peripheral joints early in the disease and ankylosis and deformity are common end results

Diagnostic Features

A Clinical Manifestations (See also the table on pages 312-313)

1. Non articular manifestations may include weakness and anorexia, fever, weight loss, edema, rash, skin nodules, myalgias, and tremors, iritis, migratory pleurisy, adenopathy, anemia, and involvement of other above mentioned body tissues.
2. The acute form of the disease is rare but may run a severe fulminating course associated with high fever, chills, cachexia, and a rapid death.
3. Mild or transient types of rheumatoid arthritis may occur.
4. Although certain joints are classically involved, any or all joints may be involved. Joint involvement may be monoarticular, but this is rare.
5. In rheumatoid arthritis of the spine (rheumatoid spondylitis), the patient may or may not be otherwise healthy but will develop recurrent low back pain associated with progressive stiffness of the spine and reduced chest expansion, often without significant involvement of the peripheral joints.

B Laboratory Data

1. Increased blood sedimentation rate and, less commonly, leukocytosis are considered to be evidence of clinical activity.
2. X-ray changes of joints and periarticular structures may be quite characteristic (see page 313) and helpful in differentiation from osteoarthritis, although osteoarthritic changes may occur coincidentally in rheumatoid arthritis and thereby confuse the picture.

Narrowing of joint spaces and ankylosis of the sacroiliac and apophyseal joints and calcification of the anterior and lateral spinal ligaments may be demonstrated in rheumatoid spondylitis.

Treatment

A General Measures

1. Rest

- a. Acute illness. Complete bed rest should be reserved for the patient with the acutely active or severe rheumatoid arthritis. Special care, including exercises, should be used to prevent deformities in bed patients, and the affected joints should be placed in the optimal functional position.

- b. Mild chronic illness. 12-hour rest periods during the daytime as well as 10-12-hour rest in bed at night are essential. Analgesics and sedative drugs (not narcotics) and physical therapy may be used judiciously to insure relaxation rest and sleep.

2. Physical activity. Carefully regulate the daily schedule of activities of the patient and allow a period for work, play, and exercise as well as for rest.

- a. Ambulatory patients. It is usually necessary to specify the hours and physical limitations for ambulatory patient according to the demands of the individual case.

- b B d p tie ts It is imperative to insti- tute a p ogram of daily systematic s to pre ent joint stiffness and m l at ophy Ref r to th s t on n physical manag m nt f arthritic joints (se p ge 323)
- 3 Diet Food ho ld be imple nou shing and p lat ble An d q te p oten and h gh vitamin diet is us ally dvis bl Since g st intestinal disord a e f q ent in rheum toid arthrit s it i often necess ry to modify the diet to tol rance C lo les should b m es ed or decr as d ding to the p t nt weight
- 4 Dietary s ppl ments
 - a Iron salts m y be indicated if anem a i p esent (s e p g 219)
 - b M l t vitamin The s of eptable m l t vitamin prep arations as g ne al health bulding m a re m y be in d ted although t i probabl that one of the vitamin has a sp if therapeuti ffe t on this on d tion V tamin D High poten y vit mun D pr parations in d ily d g ranging f om 50 000 300 000 units in d id d do a ha b n p polarized s b ng of great v lue Toxicity of the vitamin D mpounds in p long d or x i d es s d f nt and their ff ct veness has be n ques t on d by many inv t g t s Oth individual vitamin h f l d to dem nstr t ignificant b fi al re lts
- 5 Elimination of pr ip t ting facto
 - a Inf ct s E aluate the ol of syst m c fo al infec t on only as th y may apply t th individ al pati t Elumin t pe ific inf ction when v r po ible Defn t ly inf t d te th t ils etc m y be r moved o t eated as d t d It s b st to maintain a on ervat v att tud t w d the elumin tion of qu t n ble fo al unf ti s espe ially when the e tion w ll invol e extens ve m j su ge y Anti inf tiv ge ts sho ld b gi en only to comb t p lfi inf ction and not th h mat d di s p
 - b P ychog n c fa to s F quently h umat id d s ha it ons t whe th p tient is w king and living in a har ing tm pher wh he i s b j t to undu an t e h tities o s tments Imp op living hyg For co t on of u h f cto s s s tions on t physi l activity and det (abov)
- 6 P y both apy
 - a R sure pall t and r l ve e isting an tie
 - b Regul t p tients n onment t minimize mot onal diet Ke p an optim ti and ch f l att t d
 - d Expl in the n tur of th di se and th l th pati t him elf pl ys in ov ming his filn
 - e Enlist id of a t ain d pay h t t in app op late s
- 7 R li f off p n Avoid narcotics
 - a A lg i drug Give an lg ic lib ally if t l r t d t reli ve p in an aid in p eventing muscle p m and defo mity
(1) Sodium sal ylat 0.5 Gm (10 gr) (nt i oat d t p ent gast ic di t ess) v ry 2-4 hours p n p in

(2) Acetylsalicylic Acid U S P (aspirin) 0.3 to 0.6 Gm (5 to 10 gr) every 2 to 4 hours p r n pain

(3) Analgesic sedative mixture

Rx Sodium salicylate 10 to 15 giles iv

Elixir phenobarbital q s ad 120 iv

Sig 4 cc (1 dr or 1 tsp) every 4 hours p r n

- b Sedative drugs Barbiturates can be used effectively in enhancing the action of the analgesic drugs Phenobarbital 15 to 30 mg (1/4 to 1/2 gr) 3 to 4 times daily
- c Physical therapy Physical methods utilizing local heat to involved joints and proper splinting are effective in relieving pain and muscle spasm (see pages 323 to 334)
- d X ray therapy is of no value in peripheral joint involvement of rheumatoid arthritis in rheumatoid spondylitis however deep or penetrating x ray therapy carefully administered in repeated courses has proved to be of value This treatment must be administered only by trained x ray therapists

B Hormone Therapy The hormonal and steroidal agents used in the treatment of rheumatoid arthritis although they do represent a significant advance must be considered as only ancillary measures to the comprehensive approach and should probably be used only for patients who do not make satisfactory progress on more conservative treatment Perhaps the greatest disadvantage which might stem from their use aside from the very serious possibility of untoward reaction lies in the tendency of patient and physician to neglect the less spectacular but probably benefits which may be derived from general supportive treatment physical therapy and orthopedic measures These agents do not represent the long awaited specific antirheumatic factor and do not cure the disease

- 1 Corticotropin (ACTH) and the corticosteroids produce startling results in a rheumatoid arthritis but the condition regresses promptly when the drugs are discontinued Subjective improvement may appear within 6 to 48 hours after the initial dose but objective changes such as increased mobility of joints and diminished swelling occur more slowly and less constantly The period of remission following discontinuation of the steroid ranges from a few days to a few months It is mathematically predicted that for hormonal therapy in rheumatoid arthritis for the control of the acute exacerbations and the prevention of rapid progression schedules have not been established although it now appears that a satisfactory ultimate result can be obtained with smaller or more conservative doses than formerly employed Initial doses of 75 to 100 mg corticosteroids daily (orally or parenterally) should be given until control is achieved Maintenance levels of 15 to 30 mg daily may be continued indefinitely Some observer advise short periods of 1 month between the 4 to 6 week courses However the drug has been used continuously in many patients for several years without apparent harmful effect (see page 423 for further discussion of physiological dosage toxicity and of the condition)

- 2 Hydrocortisone Aetate U.S.P. aqueous suspension 10-37.5 mg intra-articularly after withdrawal of effusion fluid may be helpful for the very painful, acutely inflamed joint.

C Gold Therapy (Chrysotherapy) Although gold salts have been used extensively in rheumatoid arthritis their value remains highly controversial.

- 1 Indications Gold therapy is indicated for active rheumatoid arthritis only. Some limitations feel that it should never be used other than that it should be used only after a reasonable trial of conservative measures has failed at which other than the chance for complete remission may be better if gold therapy is used early in the disease.
- 2 Contraindications Nephritis, hepatic insufficiency, blood dyscrasias (including anemia), hemorrhagic tendencies, pregnancy, striking peripheral histology of allergy or allergies, severe diabetes mellitus, acute generalized skin disorders, ulcerative colitis, acute hematofilia, and tuberculosis.
- 3 Preparations and dosages The gold salts are usually given once a week beginning with small doses and increasing slowly. The weekly dose is increased gradually until the maximum optimal dose is being given; this amount is then continued weekly until the desired response is obtained. The maximum amount given for toxic reactions of uremia. One or two or more courses of 1000 to 2000 mg of gold salts are given with a rest of 2-6 months between courses. The value of the maintenance dose is smaller dose at regular intervals and treatment of very severe and persistent suppuration of chondrocytes, although many workers are employing this plan at present.

GOLD PREPARATIONS

Preparation	Route of Injection	Given at Once Dose Each Week		
		First Weekly Dose	Amount of Increase per Week	Optimum Maximal Weekly Dose
Gold Sodium Thio-sulfate N.F. (37% gold in quaternary solution)	I.M. or I.V.	5 mg	5 mg/week	Females 50 mg Males 75 mg
Gold Sodium Trimalate N.F. (50% gold in quaternary solution) Aurothioglucose N.F. (50% gold in suspension)	I.M. only	10 mg	Increase to 25 mg in 2nd week if tolerated increase to 50 mg in 3rd week	50 mg

- 4 Toxic reactions An average of 37% of patients (ranging in various series 8-81%) experience toxic reactions. The mortality rate is about 0.4%. The toxic effect of hyaluronidase are similar to those of other hyaluronidase not only arsenic (see page 535) and include dermatitis (mild to exfoliative), agnathous cytolysis, purpura, hepatitis, nitritoid

reactions bronchitis aplastic anemia peripheral neuritis nephritis and photosensitization

- a Reduction of frequency and severity of toxic reactions
Observe for the contraindications mentioned above. Observe patient carefully during the course of gold therapy and for a period of several weeks thereafter. Patients who are to receive gold therapy should have a complete medical examination. Before each subsequent injection ask patient how he has felt since the previous injection examine the skin and mucous membranes for dermatitis or purpura examine the urine for albumin and microscopic hematuria. Every 2 weeks obtain Hgb WBC and differential. When indicated perform special tests such as platelet counts or liver function tests. Warn patient against exposure to strong light. Withdraw drug immediately if any toxic reactions appear. Wait for a few weeks if reaction is mild and clear promptly treatment may be resumed with small doses. There is no known method of decreasing the tendency to toxicity in a given individual except perhaps through reduced dosage.
- b Treatment of toxic reactions. Withdraw drug immediately if early toxic reaction appears. Treat reactions as for allergic poisoning (see page 536). Try dimercaprol (BAL®) on all cases. (For treatment of agranulocytosis see page 231.)
- c Masked toxicity. If gold salts are used during hormonal therapy a toxic reaction may be masked appearing with explosive violence when the hormones are stopped. Therefore use gold salt with great caution during hormonal therapy.

OSTEOARTHRITIS (code No 240 912)

A chronic degenerative joint disease of undetermined cause usually of old adult life associated with varying degree of symptoms and/or disability of multiple joints. Ankylosis of joints does not take place except in the spine.

Diagnostic Features (See table on page 312-313)

- A The disease may exist with a complete absence of symptoms when symptoms are present they are usually mild.
- B Joint Symptoms Included
 - 1 Stiffness which improves with mild activity
 - 2 Aching and pain aggravated by overexertion or injury and relieved by heat rest and immobilization
 - 3 Swelling usually with joint effusion
 - 4 Deformity and malalignment of bones as a result of irregular degeneration
- C Secondary Symptoms Radiating pains occur when joint hangs in the spine due to irritation of the spinal nerve roots.

Treatment

- A General Measures Most of the general measures discussed for the treatment of rheumatoid arthritis are applicable here.

also. Emphasis must be placed upon

1. Adequate diet with total calories adjusted to meet the patient's body needs. Weight reduction is very important in obese patients to help diminish stress on joints.
2. Adequate rest and sleep. Avoidance of overfatigue is especially important.
3. Avoidance of physical activity which would cause undue trauma to joints.
4. Correct posture (page 334).

B. Drugs

1. Salicylates are indicated for the relief of pain as in the case of rheumatoid arthritis (page 315).
2. Thyroid extract may be indicated in those patients who have associated hypothyroidism.

C. For local treatment of joints see page 323. Complete rest and immobilization of involved joints for short periods may be instituted with the fear of complicating ankylosis although one must consider other harmful effects of bed rest in such patients (page 2). Hydrocortisone Acetate U.S.P. aqueous suspension 10-37.5 mg intra-articularly may be of great value in acetabular femoral joint involvement.

TABLE OF DIFFERENCES IN RESPONSE TO THERAPY

	Rheumatoid Arthritis	Osteoarthritis
Rest	Complete rest and immobilization are attended by danger of ankylosis.	Complete rest and immobilization of joints are often indicated for variable periods. Little danger of ankylosis.
Exercise	Even mild exercise produces discomfort in the acute phase of the disease.	Mild exercise causes stiffness and discomfort but undue exercise increases stiffness.
Massage (doctofuliac)	Light massage over the joints may be indicated in the convalescent or chronic disease.	Massage should be avoided directly over the bony overgrowths of the involved joints.
Chrysotherapy Peripheral joints	Often effective.	No response. Not indicated.
Surgery	No response. Not indicated.	No response. Not indicated.
X-ray therapy Peripheral joints	No response.	Some time if there is relief of pain.
Spinal	Often effective relief of pain.	Usually no response.

GONOCOCCAL ARTHRITIS (code No 24 103)

A specific infectious arthritis caused by *Neisseria gonorrhoea* (gonococcus) occurring as a secondary complication of primary infection of the genitourinary tract or conjunctivae

Diagnostic Features

History of previous genitourinary or ocular gonococcal infection and possibly of genitourinary trauma. Rheumatoid arthritis complicated by unrelated gonorrhoea occurs more commonly than gonococcal arthritis per se

A Bacteremic Phase

- 1 Fever Mild to moderate Occasionally chills
- 2 Laboratory findings
 - a Leukocytosis Mild (10 000-15 000)
 - b Blood cultures Rarely positive

B Arthritic Phase (Joint Tenderness and Bulky Involvement)

- 1 Early Evanescent polyarticular joint involvement of 3-7 days duration Joints red warm swollen and painful
- 2 Late Knees (74%) ankles (56%) feet (32%) wrists (16%) most frequently involved joints
 - a Joints initially red warm swollen and painful
 - b Ankylosis may occur in untreated cases
- 3 Laboratory findings
 - a Gonococcal complement fixation test Doubtful value especially if positive since positive complement fixation tests are known to persist many years after genital infections
 - b Cultures of synovial fluid with special culture media are the most reliable method of diagnosis but are difficult to perform

Treatment

A General Measures Several measures discussed in management of rheumatoid arthritis (page 311) and physical measures in the management of the acute phase of involvement of the various joints (page 323)

B Specific Treatment Penicillin 25 000-50 000 units I.M. every 3 hours for 7-10 days. If improvement is not apparent in 3-4 days give intrarticular injections of penicillin 10 000-20 000 units daily into the larger involved joints

BURSITIS

- | | | | |
|-------------------------|-------|-----------------|--|
| (Due to Infection) | Acute | code No 25 190) | |
| (Due to Trauma) | Acute | code No 25 4x0) | |
| (Due to Unknown Causes) | | code No 25 930) | |

Bursitis is an acute or chronic inflammation of any of the numerous bursae of the body. It may result from a local acute or chronic infection or from unknown causes. Localized pain and swelling may be observed at points around joints corresponding to anatomic bursae. Pain and limitation of motion of adjacent joints are common. Enlargement and collection of pus may be demonstrated radiologically at times.

A General Measures Analgesic (see page 32)

B Local Measures

- 1 Rest and support of involved area by lining plint bandages etc
- 2 Local heat or cold Topic applications (see page 334)
- 3 Potassium Hydrochloride U S P 0.5-2.0% injection
- 4 Hydrocortisone Acetate U S P aqueous suspension 10-37.5 mg has been reported to provide relief of acute arthritis when injected into the bursa
- 5 Aspiration of fluid from bursa Fluid should be examined
- 6 X-ray therapy in selected cases (by physician)
- 7 Surgical removal in selected cases

FIBROSITIS OR FIBROMYOSITIS

(Periarticular Fibrositis code No 24 x40)

(Chronic Myositis code No 27 190)

A large loosely defined group of acute or chronic involvements of subcutaneous tissue fibrous tissue of muscles and joint capsule of ligament tendon and fibrous connective tissue of cartilages also arises due to a wide variety of causes most of which are not conclusively determined. The condition may be manifested by pain tenderness or stiffness of any involved portion of the body. Clinical and laboratory findings are minimal or absent.

A General Measures

- 1 Eliminate aggravating factors
- 2 Rest Selection on physical management of joint disease page 324
- 3 Analgesic (see page 3)

B Local Measures

- 1 Local heat (see page 334)
- 2 Potassium Hydrochloride U S P 0.5-2.0% injection into tight areas (Of doubtful value)
- 3 X-ray therapy (by physician) in selected cases which fail to respond to other therapy
- 4 Massage and graduated exercises may be valuable
- 5 Stripping of the inflamed structure preceded by local heat or cold electrolytic procedure in fact alone may give complete relief in one treatment. If only partial relief is obtained the procedure may be repeated daily

GOUT (code No 010 741)

A disease of unknown etiology characterized by recurring acute arthritis due to deposition of sodium urate in the articular and periarticular tissue as well as in soft tissue areas throughout the body. Recurrent attacks of acute arthritis episode followed by complete asymptomatic periods is almost pathognomonic of gout and the finding of tophi is diagnostic. An elevated blood uric acid level is very common even in asymptomatic periods. X-ray evidence of punched out areas about the joints is almost diagnostic but this occurs late.

Treatment of the Acute AttacksA Specific Measures

- 1 Colchicine U S P B P is the drug of choice it should be given as early as possible in the acute attack or during the prodromata to obtain maximum benefit. Give 0.5 mg ($\frac{1}{120}$ gr) every 1 hour or 1 mg ($\frac{1}{60}$ gr) every 2 hours until there is relief from pain or until nausea or diarrhea appear then stop the drug. The usual total dose to achieve this is 4.8 mg ($\frac{1}{16}$ $\frac{1}{8}$ gr) and the pain and swelling will subside in 24-72 hours. Once the patient knows the dose that produces toxic symptoms the drug should be given in a single dose of about 1 mg ($\frac{1}{60}$ gr) less than this. Then continue colchicine 0.5 mg ($\frac{1}{120}$ gr) b i d q i d until attack has completely subsided. If diarrhea becomes too severe treat as for any acute diarrhea (see page 238).
- 2 Corticotropin (ACTH) and the cortisones provide dramatic symptomatic relief in acute episodes of gout and if given for a sufficient length of time will control most acute attacks without relapse. Since colchicine seems to be about equally effective and provides a more lasting effect it still appears to be the drug of choice. It has been observed that when corticotropin and cortisone are discontinued shortly after termination of attacks many patients promptly relapse unless colchicine is given.

B General Measures1 Drugs

- a Analgesics. At times the pain of an acute attack may be so severe that relief of pain is necessary before colchicine becomes effective. In these cases codeine with or without aspirin may be given. Morphine should be avoided for fear of addiction in this chronic disease.
- b Cinchophen or neo-cinchophen should not be used.

- 2 Rest. Bed rest is extremely important in the management of the acute attack. Bed rest should be continued for about 24 hours after the acute attack has completely subsided. Early ambulation may precipitate a recurrence.
- 3 Physical therapy is of little value during the acute attack although hot or cold compresses to the affected joints may make some patients more comfortable.

Interim Treatment

A Specific Measures Therapy limited at present to the treatment of acute attacks has been generally quite disappointing.

B General Measures

- 1 Diet. Most low purine diets (low weekly allowance of not more than avoidance of kidney, liver, sweetbread, sardines, anchovies, meat extracts) tend to become trivially inadequate and often fail to influence the hyperuricemia or course of the disease. However, it is generally thought that the restriction of high purine foods appears to be of some importance in prevention of progression of the disease. If specific foods or alcoholic beverages precipitate attacks these should be avoided. However, there is little evidence that alcohol in moderate amounts will precipitate attacks or is otherwise harmful in patients with gout.
- 2 Colchicine prophylaxis is a questionably effective measure. If it does

not influence the incidence of attacks which should be relieved of the paroxysms

Treatment of Complications

A Chronic Gouty Arthritis In recent years the outlook for patients with this disease has greatly improved. In many cases the progress of the disease is arrested and in many cases the absorption of gouty deposits may occur. This condition is best treated by a low purine diet and the newer uricosuric drugs.

1 Uricuric drugs

a Probenecid N N D (Benemid) agent which blocks the tubular reabsorption of filtered urate has been proved to relieve chronic gouty arthritis. Dose of 0.5 Gm (7½ gr) b.i.d. over long periods. It may be given indefinitely if tophi are visible. The blood uric acid is greater than 7 mg% if it keeps a few weeks. The uric acid should be kept alkaline with sodium bicarbonate or sodium citrate and fluid intake should be high. Acute episodes of gout may occasionally be precipitated by this treatment but this is diminished if the administration is continued. Full dose of colchicine may be given with Benemid.

b Salicylate. Large doses of all salicylates up to 5 Gm (75 gr) daily have been reported to produce uricosuric effect similar to the above with relief of symptoms. Do not combine directly with Benemid. Phenylbutazone N N D (Butazolidin®) has been reported to provide relief of acute inflammation and pain by lowering of serum uric acid but this is a relatively dangerous technique especially when used for long periods. It diminishes daily to 100 mg tablets 3 tablets daily as needed. To therapy should be helpful at times preventing further deposition of uric acid.

2 Sulfur may offer some help in relieving mechanical deformities but is not always effective.

B Renal Complication The formation of uric acid in the kidney is decreased if patients are encouraged to drink at least 3000 cc of fluid daily. On alkaline fluid little can be done to dissolve the stones although forcing fluids and alkalizing the urine with 8-16 Gm (4 dr) of sodium citrate per day may be helpful at times preventing further stone formation.

PHYSICAL MANAGEMENT OF ARTHRITIC JOINTS (PHYSICAL THERAPY)

General Principles

Certain general principles which apply to the treatment of diseased joints are emphasized.

1 Arrange posture of affected joints in comfortable position which will provide for optimal physical use in the event that joint motion has been equally lost.

2 In the ankylosing form of arthritis after the acute process

3 4 Physi al Therapy

has subsided employ careful active exercises or passive mobility at once early and regularly as tolerated in order to prevent deformity and to preserve joint motion

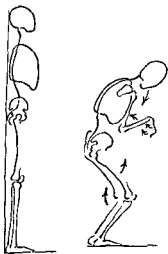
3 Avoid measures which cause a persistent increase in symptoms. So called routine measures e.g. heat are not uniformly tolerated by patients. The correct balance of heat massage rest and exercise must be planned for each patient.

4 Patients with joint disease (particularly rheumatoid or suppurative arthritis) are in constant threat of deformity. Guard particularly against flexion deformities.

5 The services of a specialist in physical therapy should be utilized whenever possible.

6 If the arthritis is severe the course of the disease seems unfavorable or ankylosis appears inevitable early consultation with a specialist is imperative. Special orthopedic measures such as manipulation under anesthesia, traction, special casts, braces and corsets and surgical measures including arthroplasty, arthrodesis, tenotomy, arthrodesis and synovectomy may be required.

7 Emphasize to the patient the importance of complete cooperation and his responsibility with the physical therapy program at home as well as in the office or hospital. Stress the importance of year-round continuation of treatment if necessary. Instruct the patient and/or his family and friends as to the extent and proper use of heat, immobilization and passive mobilization at home conditions.



Normal Posture and the Deforming Tendencies of Joint Disease

OPTIMAL FUNCTIONAL POSITION OF JOINTS

Should ankylosis seem inevitable despite adequate therapy or desirable early orthopedic consultation is imperative. The following table gives certain commonly accepted optimal functional positions in which joints may be permitted to fuse. It must be emphasized however that a given position is functional for a given individual depending upon such factors as living habits, occupation, recreation, and personal preferences.

Joint	Position
Shoulder	Arm abducted at about 75° the elbow joint flexed in line with the anterior chest and with partial pronation of the forearm
Elbow	Elbow at slightly less than 90° with the forearm in a position midway between supination and pronation. Exceptions: laborers full supination, clerical work full pronation.
Wrist	Slight (30°) dorsiflexion
Hand	Partial flexion of fingers and metacarpophalangeal joints and partial apposition of thumb
Hip	Unilateral in lying: Extension with minimal abduction and minimal lateral rotation. Bilateral in sitting: Slight flexion of one hip joint and the other as above
Knee	Active individuals: Full extension. Sedentary individuals: 30° of flexion. Young individuals with active epiphyses: Only slight flexion
Ankle	Foot at 90° angle with leg minimally everted

REST, IMMOBILIZATION AND SUPPORT

Spine Rest in a comfortable position on a flat firm bed without pillow as described essential. A 3 x 5 ft. plywood or other simple type of bedboard placed under the thin firm hair or felt mattress is advisable. Immobilization and support of the spine by a simple well applied adhesive strapping may give prompt temporary relief. Suitable corset or back braces may be employed on ambulatory patients with mild symptoms. Special body molds (plaster shell) or rigid jackets may be advised by the orthopedist for patients confined to bed.

Cervical Spine Rest in a comfortable position on a flat firm bed without a pillow (as above). Immobilization and support of head may be accomplished by special orthopedic or home made collars. The latter may be made simply by folding a soft bath towel twice lengthwise and wrapping snugly around neck and fastening with pins. Traction may be necessary if deformity is present or pain is severe.

Shoulder Rest in bed in a comfortable position on a flat firm bed with use of a pillow (as above). Support the arm with pillows in a position of intermediate abduction and external rotation. After

patient is ambulatory the arm of the involved shoulder may be supported by a sling

Elbow Support the arm and hand (thumb and fingers free) in a molded bivalve plaster cast with the elbow in a position of maximum tolerated extension. This is to combat the natural flexion tendency.

Wrist Provide rest and support for the hand in a bivalve splint which corrects the natural deforming tendency toward flexion and ulnar deviation. At first the splint should be worn continuously except for removal 1 or 2 times daily to permit physical therapy. Later the splint need be worn only in bed.

Hand A supporting plaster splint fitted into the palm and extending to form a pocket for the partially flexed fingers may prevent the natural deforming tendency of hyperextension of the metacarpophalangeal and distal interphalangeal joints and flexion of the proximal interphalangeal joints. This splint should be removed 2 to 3 times daily to permit physical therapy.

Hip The patient should be able to bear weight. A detachable plaster hip spica may be used to provide support and rest for the acutely involved hip joint. It may be worn all night but it must be removed at least 2 or 3 times daily to permit physical therapy (at least in the case of rheumatoid joints). The patient is instructed to lie prone in bed with 1 or 2 pillows (as necessary) under the abdomen to flex the hips for $\frac{1}{2}$ to 1 hour 1 or 2 times daily. (The weight of the body is utilized as a load against the powerful flexor muscles of the thighs.) The pillows may be removed as tolerated and a thigh flexion deformity is corrected.

Knees The patient should initially be able to bear weight. Bearing up on the acute joint should be restricted or prohibited. In mild and nondeforming joint the posture of plaster splint is of interest to us and will suffice. It should be worn almost continuously while the patient is in bed particularly during the night. Adjustable splints with slight flexion may be employed on patient who is able to walk. When joint involvement is more marked a definite deformity is present correct plaster cast applied in position of maximum correction and left off 2 days. The cast is bivalved and removed for physical therapy twice a day. New casts are made to provide further correction as indicated. During convalescent or chronic phase provide support for the knee with elastic bandage posterior splint or special orthopedic brace.

Ankle and Foot Weight bearing up on the acutely involved joint must be prohibited. Provide a cradle or large pillow at the foot of the bed in order to hold the bedclothes off the feet. A supporting removable plaster boot cast (with tips of toes exposed) is valuable. Adjustable or ritual bivalved plaster foot cast may be employed for the gradual correction of deformities. Provide well built shoes allowing proper length and width for toes stability and a suitable arch support (sponge rubber or felt pad a quilt satisfactory). Correct abnormalities and deformities of the hip and knee joints which produce mechanical strain on the feet.

Chapter 13

TECHNIQUES OF MEDICAL REHABILITATION

By rehabilitation is meant the restoration of the physically handicapped patient to his maximum physical and social capacity. To be most effective it should be initiated at the onset of illness, be it acute or chronic. Difficult therapy and rehabilitation is often difficult to achieve in which case appropriate compensation must be made in order to render the best possible service to the patient.

Reduced mobility is often caused by the original illness but by inactivity (e.g. bed rest) and atrophy in a faulty position because of weakness or pain. For example, fixation on trauma, dislocations and painful shoulders in hemiplegia and myocardial infarction are avoidable if rehabilitation is started early. Furthermore, the patient's psychological response to his illness is much more favorable if he knows that the physical goal in treatment goes beyond the immediate medical problem to include restoration to useful function.

The techniques used in medical rehabilitation are primarily physical and educational. Physical measures consist of positioning, splinting, bracing and exercise. Within the limits of residual ability the patient must be trained to maximize function with or without the aid of orthopedic devices. Rehabilitation procedures are aided with the help of nursing staff and therapists (physical, occupational speech) under the direction of the physician. In the home management of a disabled patient members of the family may be instructed in rehabilitation procedures.

PREVENTION OF DISUSE PHENOMENA

Immobilization and inactivity of a part of the body leads to changes which may be grouped under the term disuse phenomena (etiology page 328). Lack of mobility may be caused by paralysis, pain, limited joint motion (ankylosis), contracture, abscess, amputations, arthrolysis and restricted activity of medical or psychological causes. Disuse phenomena lead to further disability and thereby aggravation and extension of disuse.

The main reasons for the prevention of disuse phenomena are as follows:

- (1) Prevention of joint motion (page 334-2)
- (2) Proper positioning and support (page 325)
- (3) Changing position (page 329) including the tilt table (page 330)
- (4) Exercises (page 334) (See also B d E page 329 Stand up Exercises page 331 and Occupational Therapy page 334-2)

Cause and Prevention of Disuse Phenomena

	Cause	Prevention
Muscle atrophy (weak)	Lack of exercise	Exercise
Joint contracture (limited range)	Lack of joint motion	Change position passively range of motion splinting and support
Orthostatic hypotension	Lack of tilt position	Tilt table and stand up slowly
Bone atrophy (osteoporosis and urinary lithiasis)	Lack of weight bearing and mobilization	Tilt table and stand passively 1-2 times a day
Deubitus ulcers	Prolonged pressure	Change position frequently
Venous thrombosis	Stagnation of blood flow	Change position frequently
Hypertension	Lack of chest expansion	Change position frequently
Urinary incontinence	Unreliability of external sphincter at time of voiding	Urinal bedpan used while doing voiding exercises

CHANGE OF POSITION

Frequent change of position is the most important single measure in the management of the disabled patient. Patient susceptible to pressure sores require frequent shifting. The time in each position should be prescribed by the physician in the form of a precise schedule including all therapeutic and nursing procedures. Routine orders such as up in chair as tolerated are ineffective and may even be harmful. For example, a paraplegic who tolerates six hours sitting in a chair will develop bilateral ischial pressure sores.

The following is a sample change of position schedule which was prescribed for a patient with a severe neurologic disease (early transverse spinal cord lesion). Let us assume that because of flaccid paralysis sensory loss, heavy weight, and lack of substantial fat padding, the maximum time for pressure over the sacrum, trochanters, and ischia must be limited to one-half hour.

Sample Change of Position Schedule

7:00-7:30 a.m.	Bed bath	7:30-8:30 p.m.	Tilt table
7:30-8:00 a.m.	Chair (backrest)	8:30-9:30 p.m.	Bed (half hr rt)
8:00-9:00 a.m.	Bed (half hr rt)		1 ft 1 ft deubitus)
	1 ft and half h 1 ft 1 ft	4:30-5:00 p.m.	Chair (dinner)
	deubitus)	5:00-7:00 p.m.	Bed (half hr rt)
9:00-9:30 a.m.	Commence (bowel training)		1 ft deubitus half hr Fowler's
9:30-10:30 a.m.	Bed (half h)		half h 1 ft lat deubitus
	pine half hr Fowler's position)		half h pi)
10:30-11:00 a.m.	Tilt table	7:00-10:00 p.m.	Bed (position)
11:00 a.m.-12:00 noon	Bed (position)	10:00-11:30 p.m.	Bed (half hr rt)
12:00 noon-12:30 p.m.	Chair		1 ft deubitus half h pi)
	(lunch)		half h 1 ft 1 ft deubitus)
12:30-3:00 p.m.	Bed (half h rt)	11:30 p.m.-2:30 a.m.	Bed (position)
	1 ft deubitus half h spine	2:30-4:00 a.m.	Bed (half h rt)
	half h 1 ft lat deubitus		1 ft deubitus half h pi)
	on hr position)		half h 1 ft 1 ft deubitus)
		4:00-7:00 a.m.	Bed (position)

In a patient in prolonged coma the need for change of position is similar but the chair and tilt table have to be eliminated in favor of the prone position. The prone position is well tolerated by the comatose patient and can be maintained for several hours without causing pressure sores and also will permit postural drainage of respiratory secretions. In an elderly patient in danger of orthostatic hypotension however the chair and tilt table may dominate the day schedule. A chair period should not exceed one hour in the ideal patient because of the danger of knee and hip flexion contractures.

While the comatose and severely paralyzed patient depends entirely on the nursing staff for his changing of position the patient with some physical ability should carry out his change of position by himself or with minor assistance. (See Exercises below.)

BED AND BED EXERCISES

To permit change of position as well as getting in and out the bed must have the following features: (1) A firm mattress (preferably two inch foam rubber over a wooden board). (2) A siderail if the patient cannot turn without holding on. (3) An overhead frame with trapze for sitting up and getting out of bed. (4) A footboard movable to prevent the sheet from pressing on the feet. In the paraplegic the feet should rest again in the footboard to prevent foot drop. (5) The height of the bed should be the tilt table height which is height according to the need of the patient. (6) The bed must be secure against the floor so that it does not slide or roll when the patient gets out or in or leans against it.

Bed Exercises

Bed exercises comprise the groups of actively active change of position, self care and occupational therapy and special exercises.

A Active Change of Position. This may be carried out by the patient under the direction of the nurse or the physician as necessary with some assistance. Example. To turn from supine to the right side move the body toward the left edge of the bed pull up the left knee each with the left arm for the right side rail and pull the body over to the right side.

B Self-care. Feeding, washing, wiping, etc. should be done by the patient as much as possible unless contraindicated.

C Special Bed Exercises

- 1 Sit straight up (Bed flat patient supine neck red heads on mattress side rails) Raise right leg to vertical position with knee extended. Lower slowly onto bed. Do the same with the left leg. Repeat each leg ten times.
- 2 Sit up (Head of bed raised to 45 degrees patient pin on neck red mid-reclining hands folded behind neck legs straight) Raise trunk into sitting position. Recline slowly. Repeat ten times.
- 3 Chinings (Bed flat patient prone both hands on overhead trapeze) Raise trunk by bending elbow until shoulders are at the level of the trapeze. Lower trunk slowly. Repeat ten times.

CONTRAINDICATIONS When complete bed rest is indicated (see page 1) all bed exercises are contraindicated. Certain of the exercises may be contraindicated because of local condition (pain

fracture etc.) The amount of activity should often be gradually increased, and the exercises prescribed must be chosen accordingly.

TILT TABLE

The tilt table is a useful device for the gradual establishment of weight bearing or erect position and for retraining in weight bearing or erect position in patients unable to stand. Weight bearing helps prevent or counteracts osteoporosis in the lower extremities. The erect position serves as a change of pressure area prevents flexion contractures develops extensor spasticity in upper motor neuron lesions facilitates chest expansion and urinary drainage and prevents loss of sense of verticality and development of orthostatic hypotension. Gradual establishment of erect position helps overcome existing orthostatic hypotension. The tilt table is contraindicated in unconsciousness severe pain and acute illness.

Technique

A tilt table resembles a stretcher whose top can be tilted gradually from a horizontal to a vertical position. A footboard at the lower end prevents the patient from sliding down and allows weight bearing. The patient may be strapped to the table top across the pelvis knees and chest as needed. He should not be immobilized more than necessary and should perform some exercise activity during the standing period. The angle of tilt is determined by the desired amount of weight bearing and erect position. It is usually between 45 and 80°.

In most patients for whom the tilt table is indicated, i.e., patients with paralysis who have been in bed for a long period, the tilted position must be established gradually over a period of days and weeks. Once the patient is used to the tilt table, he may remain in the maximum tilt position up to one hour two or three times daily. Tilt table treatment is begun as follows:

1. Bind the patient's lower extremities tightly with elastic bandages and apply a tight abdominal binder (asci-tetus).
2. Place the patient on a flat tilt table and fasten the straps across the pelvis knees and chest.
3. Apply a BP cuff to the arm and record the BP.
4. Tilt the table 15° and take BP every two or three minutes.
 - a. If the patient feels faint or dizzy and the BP drops, return to flat position and start over raising the table only 10° (If the patient complains of dizziness but BP does not drop, encourage him to stay up and proceed as below.)
 - b. If the patient feels all right but the BP drops, watch closely. After a few minutes the BP may rise and stabilize slightly below the patient's normal pressure. Do not increase tilt but maintain a 15° tilt for 15-30 minutes. If the BP continues to drop, the patient will soon feel dizzy and should be returned to the flat position.
 - c. If the patient feels all right and the BP maintains itself for several minutes, increase tilt by another 15°.
5. Continue procedure until the patient tolerates the desired tilt. Then gradually eliminate bandages and abdominal binder.

STAND-UP EXERCISES

Stand p e e cises are indi t d w h n th is a d for w ght bearing nd e e t position (see Tilt Tabl p g 330) at ngth ing of calf th gh and t nk mu cl to in a e spirat ry and circu l tory eff cy to de elop tanding bala and in prep ration for ambul tion unl ss combin d w th stand up exercises) The e exe cises ar ontr indicated in ac te ill e diac d omp sation and e rly myo a d l inf ctio

Techn

Th i e consists of standi g up from a chair ing the p we of th kn hip and t unk exte n The r s can b do on o e leg If th patn t is y weak a v ry high chair m st be us d Ac d g to th h ght and str ngth of the p tient wooden bl cks six fo two i che high with a six by six in h ba e and h ll wed c nt s are pla d unde the leg f th h Th p ti t f ces th foot end of th b d and m y teady him lf by holding o to th b d f me He m st w ar sho s which do n t slip on th floo A two by tw l bo d bra ed g inst the legs of the bed is u ef l m tabili g the pat nt foot Th p t n of the w ight b i g l g s s h that th heel lines up with th fr t l g s of th cha The umber and sp d f t nd p d pe d upon the indi vidual p t nt Two to ix ion a d y a e u ally pr s b d best at the b g nni g and/o at th nd of th h i pe id An v g p escription is t n tand up pe e ion t the rat of two p r min t

REHABILITATION OF THE HEMIPLEGIC PATIENT

Advan i phys cal ru d i h egl n whop to the p ti nt wh suff s f m h miple g a a diti n which is ounte ed mor a d m in clin cal m dici The followi g p gram is in t d d to ser nly guide it appl s to th typ cal cas f er bral l accide t b t th p i ciples ar th me in h miple g a of a y tiol gy

B d Ph

St ts on o d or th d d y f illness o as soon a the patie t is on s Th patient s bed sho ld b f h i h ight and h old h e aid rails and an ov rh ad t pe e

- A E is Start w th t minut s of i e e ery two h and inc s to 30 m nut of x i e e y tw ho s
- 1 With good arm a d leg turn from b k to ide to abd me then to oth ide a d th n b k Repeat in ppo ite dir to
 - 2 W th good h nd o t pe e pull t s t t ng po ition and back
 - 3 M e sid w y pw rds and d wnw ds in b d
 - 4 Sit p on edge of bed with ide rail moved legs da gling and m e along dg of b d with aid of good a m a d l g
- B S If C (all d with g od hand)
- 1 T llet activities Wash f a d h ds comb hai sh ve
 - 2 Fe ding ti it s At fl t i b d with b ck oll d p l t r itting o edg of bed
- C B i g N d g bed ph s

Standing Phase

Starts three to five days after beginning bed phase replaces bed phase as soon as possible. Patient is placed in a chair with his good side next to the foot of the bed the vertical bar of the overhead frame in reach of his good hand and the paralyzed arm in a sling.

A Exercise (See also page 331) Start with ten minutes of exercise every two hours and increase to 30 minutes every two hours.

- 1 Raise to standing position on good leg. Sit back.
- 2 Standing with good hand on vertical bar of overhead frame perform slight knee bend and straighten up. Repeat with gradually deeper knee bends.
- 3 Stand with good hand on vertical bar of bed frame. Go up on toes come back down.

B Self Care (using good hand)

- 1 Toilet activities. Complete bath in bed.
- 2 Dressing activities. Dress and undress except for shoes.

C Bracing

- 1 Fit wooden splint (attached to volar surface with ace bandage or straps) from one inch below the elbow to one half inch beyond the fingertips of the paralyzed arm.
- 2 Keep paralyzed arm in sling to prevent pull on shoulder.
- 3 If after two weeks the paralyzed leg still remains completely flail a long leg brace is needed in order to continue rehabilitation.

Stair climbing Phase

Starts two to ten days after the beginning of the standing phase and should replace standing phase as soon as possible.

A Exercise Performed four times a day increasing from several steps to a whole flight of stairs. The patient is placed in a chair facing the foot of a flight of stairs the good arm next to the banister. The paralyzed arm is splinted and in a sling and the paralyzed leg is in a long leg brace if needed.

- 1 Pull to standing position holding to the banister with the good hand step up one step with the good leg then pull paralyzed leg up to the same step. Continue for several steps.
- 2 Step backward and down with the paralyzed leg and put the good leg down next to it. Continue for several steps.
- 3 While several stairs up turn toward and reach over to the opposite banister. Step forward and down with the paralyzed leg. Then place good leg next to paralyzed leg and continue.

B Self Care Complete toilet and feeding and dressing activities should be possible by this time.

C Bracing

- 1 Long leg brace if indicated (see Standing Phase).
- 2 If patient has a foot drop during stair climbing he should have a short leg brace with a 90° posterior stop at ankle.
- 3 If the patient shows evidence of inversion or eversion of the foot he should have a short leg brace with a T strap.
- 4 If function has returned to the paralyzed hand the splint may be discarded. Otherwise it should be worn intermittently.

Cane walking Phase

Starts as soon as the patient is capable of walking up and down a whole flight of stairs without tiring. Paralyzed arm is kept in a sling and cane is held with good hand. Two different types of

cane are recommended for the hemiplegic patient

A Single (For fearful patients or patients with poor balance)
Move cane forward, place good foot next to cane and then drag paralyzed foot next to good foot

B Four Standing on good leg place cane and paralyzed leg forward simultaneously and put weight on them. Swing good leg through in front of cane and paralyzed leg and put weight on it. Continue in this fashion

Special Problems in Hemiplegic Patients

A Care of the Paralyzed Upper Extremity

- 1 Complete absence of function. In most cases, no useful function returns to the paralyzed upper extremity and the wrist and hand are best supported in the plant. The sling may be discarded later when the shoulder muscles become spastic and the patient feels limited by the sling. With his good hand the patient should move the paralyzed fingers, wrist and elbow through the full range of motion twice a day in order to move the paralyzed shoulder through the full range of motion. The patient may need a good thigh and/or shoulder brace by means of which the paralyzed arm (tied at the wrist) can be pulled up as high as possible with the good arm.
- 2 Partial function. If only partial function returns to the paralyzed extremity the patient should use it only for the extent to which it is helpful or expedient. For other activities the patient should be trained in the use of the good extremity.
- 3 Complete function. If complete function returns the patient should use the extremity as much as possible.

B Treatment of Aphasia. If aphasia occurs, perhaps the apy (differently in the period) should be started as soon as possible. If sensory or receptive aphasia is present, the above program may be rendered extremely difficult since it is based on the ability of the patient to understand what is required of him.

C Care of Homonymous (minor problem). If homonymous is present the patient should be trained to turn his head to the homonymous side in order to bring his visual field in front of him. Later, the adjustment in the visual field occurs.

D Care of Spontaneous. Some hemiplegics are incontinent in the early phase. A indwelling catheter is rarely necessary. The patient should be reminded to empty his bladder voluntarily at hourly intervals. The interval can be gradually increased.

E Organic Mental Syndrome. When this is present, the whole rehabilitation program becomes difficult. The patient may either understand or may be unable to understand. The confusion may be present at one time and absent at another and adjustment should be taken of the patient's individual needs. Organic mental syndrome occurs usually in patients who have had a cerebral stroke. The patient's mental state usually improves considerably during an active rehabilitation program.

REHABILITATION THERAPY

Physical Therapy

Physical therapy can be defined as a method of treatment and prevention of musculoskeletal disorders by the use of light

electricity cold heat exercise and passive mobilization. The latter three are used most commonly.

- A Heat Heat therapy is indicated in acute and convalescent diseases of the joint muscles fasciae tendons and bursae to relieve pain and to reduce muscle spasm and for chronic involvement of these areas to relieve pain reduce muscle spasm hasten recovery and to serve as an adjunct or preparation for other physical therapy methods. It is contraindicated in local diseases of the skin peripheral vascular disease (as circulatory insufficiency see page 207) and in patients with loss of sensation.

Technic Place the part to be treated in a comfortable and relaxed position. Begin slowly and cautiously. Treat for short periods not longer than 15 to 20 minutes initially. When the skin is pink and moist discontinue. Start with lower temperatures adjusted to patient's tolerance. Gradually increase time and temperature as tolerated and indicated (average time is 30 minutes). Avoid drafts. Following treatments provide protective covering for 20 to 30 minutes to avoid chilling.

There is no evidence that any of the specific methods given below have more therapeutic value than others or are preferable in certain conditions although they differ in penetration.

- 1 Conductive heat (direct contact) Hot water bottle or electric heating pad. Moist heat can be administered to local areas as hot compresses (hot packs) soaks or whirlpool or paraffin baths. Tub baths and the Hubbard tank permit submersion of the whole body and simultaneous administration of stretching and exercise (see below).
- 2 Radiant heat Electric bulbs or infra red lamps. A baker's salamander reflecting hood with several bulbs.
- 3 Conversive heat Heat may be developed by the resistance of tissues to the passage of high frequency wave or ultrasonic vibrations. Treatment of this sort may be more effective than other forms of heat treatment in certain patients with involvement of the spine and large joints. It is contraindicated in late pregnancy and in patient with malignancy. Long wave diathermy short wave diathermy militherm (adar wave lengths) and ultrasound equipment are used for this purpose and certain points deserve consideration.
 - a Special equipment is required and treatments must be administered by a trained operator. Never permit self treatment with diathermy by a patient at home.
 - b Rarely case titles since other simple methods are usually equally effective.
 - c No specific therapeutic effects other than heating action.

B Exercise

- 1 Therapeutic exercise is a voluntary active motion of part or all of the body designed to produce improvement of function.
 - a Assisted exercise Voluntary movement by the patient directed and assisted by the therapist.
 - b Independent exercise Voluntary movement under direction but without assistance.
 - c Resistive exercise Voluntary movement against graded loads or resistance as the final step in the repetitive exercise. Specific remedial exercises and occupational therapy represent a more advanced type of resistive exercise.

- 2 Specific planned remedial exercises. A great many specific functional exercises have been designed to correct abnormality of joint function and muscle weakness. The various home remedial investigations and remedies described may be useful but must always be modified to meet individual need. Some of the common and well accepted exercises include (1) weight and pulley exercises for elbow and shoulder girdle joint (2) hand exercises (squeezing rubber ball or sponge or hand pump) of fist making exercises (may be performed under water in basin or tub) (3) drawbar exercises (polio Hobbatake) for pin and girdle palsy and lower extremities (4) parallel support devices to permit weight bearing of upper lower extremity joints and (5) postural exercises. Postural exercises are carried out as follows

Sitting. Sit in chair and maintain tall upright position for 2-5 minutes.

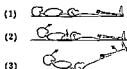
b Standing (see below)

- (1) Stand against wall with knees perfectly flexed
- (2) Relax the cervical and lumbar region of the spine and pinch the spine together
- (3) Gradually tighten knees

(1) (2) (3)



Standing



Lying Down

Lying down (see below)

- (1) Lie back
- (2) Pinch the chest and upper lumbar spine and knees to floor
- (3) Relax the legs and chest simultaneously

d Walking

- (1) Walk with weight evenly distributed and feet pointed straight ahead
- (2) Roll the hips and buttocks together by contracting the buttocks and abdominal muscles, pinch by fitting the pelvis back to straighten the limb
- (3) Relax the chest, the elbow, the thigh, the thorax, the wrist and the hand
- (4) Stretch the back part of the foot of the heel toward the lung the straight thigh the wrist and the hand
- (5) Start to walk till possible

C Passive Mobilization The principal types of passive motion exercises are described below. Stretching and range of motion exercises may be prescribed as often as every 2 hours. Massage is usually given once a day; manipulation not often than once or twice a week. The minimum of therapy depends upon the patient's condition. Passive motion and stretching should be repeated several times for each joint. If only one joint is involved the procedure will last only 2 to 3 minutes. If several are treated it will last longer but should not exceed 15 minutes. If all joints cannot be treated in one session the number of treatments should be increased and the joints alternated.

1. Passive motion of joint through the existing range. This is done by a physical therapist or other person or by the patient himself. The movement is slow; it should cover the complete available range and be repeated several times. No force should be used and the patient should be relaxed. The objective is to prevent loss of range of motion, particularly in patients immobilized by splints, slings, and so on, etc.
2. Stretching. Similar to passive motion and somewhat forceful, it is carried slightly beyond the existing range. This procedure may cause a certain amount of pain and should be preceded by analgesics. Stretching should be performed only by a physical therapist. Indication. In defining types of arthritis, stretching is used to decrease flexion contractures. In osteoarthritis and fibrositis it may actually lead to cure by giving complete pain relief. In postural correction stretching is an adjunct to active exercises.
3. Manipulation. A passive rather than a forced mobilization of joint in a direction which is not used physiologically, e.g., sideward motion or rotation in a metacarpophalangeal joint. Manipulation should be done only by a physician, preferably with anesthesia. It is used to break up painful intra-articular adhesions. It may also be used instead of stretching. It has the advantage that the mobilization cannot be counteracted by voluntary contraction or spasm because of the direction of the pull.
4. Massage (minor physiological value). Essentially a mobilization of soft tissue by direct manual or digital action.

OCCUPATIONAL THERAPY

Occupational therapy is defined as a medically prescribed activity with the emphasis on purposeful activities. These include (1) self-care training (e.g., feeding, washing, dressing), (2) recreational activities (e.g., weaving, wood carving, needlework, painting), and (3) provocative activities (e.g., typing, business machine operation, machine hop work).

SPEECH THERAPY

Speech therapy is the medically prescribed correction of speech disturbances resulting from (1) psychiatric causes (e.g., stuttering), (2) neurological causes (e.g., aphasia, vocal cord paralysis), or (3) otolaryngological causes (e.g., cleft palate, deafness).

Chapter 14

DISEASES OF THE NERVOUS SYSTEM

DISORDERS OF CONSCIOUSNESS

Disturbance of the consciousness may be associated with decreased mental activity (e.g. stupor or morose) or increased mental activity (e.g. excitement delirium mania). Solid turban may suggest complete obliteration of consciousness. The patient's reaction to the stimulus depends upon the nature and intensity of the stimulus and the physiological and mental status of the individual. Some of the aetiological factors are trauma, cerebral abscesses, drug intoxication, poisoning, infectious diseases, meningitis, overwhelming infection, convulsions, disorders and cardiac decompensation.

STUPOR (code No. 933) and COMA (code No. 932)

Stupor ranges from partial to almost complete loss of consciousness. Complete unconsciousness is from which patient cannot be aroused voluntarily. Mental functions are lost.

Diagnosis

A. History - It is to be obtained during the interview. Valuable information may also be obtained from the patient's relatives and attendants. Inquire carefully about the patient's past medical history, physical, mental, emotional illness, diabetes, trauma, alcohol, drug, epilepsy or hypertenension.

B. Physical Examination - Place particular emphasis on vital signs, deep reflexes, extensor reflexes and neurological examination. Do not use solid turban as a total hindrance. Clean the head thoroughly.

C. Laboratory Procedures

1. Check the patient if necessary and examine urine particularly for blood, sugar and acetone.
2. Take Hgb, WBC, differential count and haematocrit.
3. Draw blood for NPN, glucose, blood urea nitrogen and send for diagnosis of anemia, diabetic coma and hypotension.
4. Lumbar puncture should be considered if all other points are negative.
5. Special studies may be indicated, e.g. blood chemistry and analysis of body fluids for various toxins.
6. Skull x-ray when indicated.

Treatment

A Emergency Measures *Maintain life until specific diagnosis is made and treatment administered*

- 1 Maintain adequate respiration First determine the cause of any respiratory difficulty (e.g. obstruction pulmonary disease depression of respiratory center vascular collapse)
 - a *Keep airways open* Place patient on his side or abdomen with face to the side and head well extended (NEVER on his back or with head flexed) If necessary pull tongue forward with fingers or forceps and maintain in an extended position (e.g. by pharyngeal airway) Aspirate mucus blood and saliva from the mouth and nose with a lubricated soft rubber catheter If no suction apparatus is available use a 25-50 cc syringe Endotracheal catheterization or tracheotomy may be necessary (CAUTION Do not allow endotracheal tube to remain for more than 2 hours because of danger of laryngeal edema and further obstruction upon its removal) The services of a trained anesthesiologist or otolaryngologist are desirable for this
 - b Artificial respiration may be administered if respiration has ceased or is failing (see page 150)
 - c Oxygen may be administered by mask catheter or tent as indicated (see page 138)
- 2 *SHOCK* Institute immediate treatment if patient is in shock or may suffer shock (see page 27)

B General Measures

- 1 Constant observation of the patient must be maintained
- 2 Unless contraindicated place in shock position (see page 2) and change body positions every 1/2 hour to prevent hypostatic pneumonia and skin ulcerations
- 3 Catheterize patient if urination persists for longer than 8 to 12 hours and patient fails to void If necessary insert an indwelling catheter Use sterile technique
- 4 Nutrition and hydration Provide proper fluid and nutrition by I.V. glucose amino acids and saline solutions (see page 20) for the first few days until the patient is able to take fluids by mouth If the patient is comatose for more than 2-3 days tube feedings must be employed (see page 59)
- 5 Sedation
 - a Whenever possible avoid sedation or other medication until a specific diagnosis has been made
 - b Sedation with paraldehyde or barbiturates may be necessary for mild restlessness in those cases not due to barbiturate or other drug toxicity

C Specific Measures Treat specific cause such as fevers infections toxin (see specific diseases)

DELIRIUM (code No. 931) and MANIA (code No. 037)

Delirium is characterized by mental disturbances (e.g. illusions delusions and hallucinations) physical excitement with restlessness and lack of coherence

Mania is often temporary characterized by wild exaltation and at times by illusions delusions and hallucinatory trends

These two conditions are discussed together because they have many points in common. The principal therapeutic differences are in the choice of sedative and hypnotic medications. Although most sedative and hypnotic drug in proper dosage may be used with relative impunity in many, the number of drugs which can be employed is limited. Chloral hydrate is contraindicated in actual alcohol delirium. (For diagnosis see Comp p 335.)

Treatment

A. Preotect From Physiological Injury

1. Quarter Use safest room available preferably on lowest floor of building
2. Windows Screened or otherwise protect windows. Locked heavy screens are most desirable.
3. Furniture Remove all furniture and furnishing from the room except a low bed and bed with side board or at times simply a mattress on the floor. The room must be free of sharp objects.
4. Avoid mechanical irritants whenever possible except for specific medical or surgical reasons. Use chemical restraint to use hydrotherapy as mentioned later. Observe for suicidal or destructive tendencies.

B. Rest Patient

1. Be kindly and understanding. Reassure patient's actions as those of a confused and sick person.
2. Lighting and noises. See that the room is adequately lighted both day and night and free from shadow. Unusual noise should be avoided but familiar sounds may actually serve to ease the patient. Remember that the patient may be confused and will misinterpret strange sensory stimuli.
3. Help the patient to understand what is happening and why he is in his particular situation. Do not misinterpret. Explain diagnosis and therapeutic procedure when necessary.
4. Relatives and friends. Recruit aid from relatives and friends. Encourage family figures they may serve to lessen the patient's apprehension. However, psychiatric patients frequently become disturbed under these circumstances.
5. Constant nurse attendance is necessary.

C. Sedative and Hypnotic Drug Treatment

1. Chlorpromazine Hydrochloride U.S.P. (Thorazine®)
 - a. In acutely agitated or disturbed patient, chlorpromazine may be used effectively to relieve agitation. The initial dose is 25-50 mg ($\frac{3}{8}$ - $\frac{3}{4}$ gr) by deep I.M. injection. Subsequent injection may be required at intervals of 4 to 6 hours. In less acutely disturbed patients give oral haloperidol 25 mg ($\frac{3}{8}$ gr) tid in response if necessary to 100-400 mg ($1\frac{1}{2}$ - 6 gr) daily.
 - b. Toxicity. Jaundice is noted occasionally (about 1-5%) but is reversible with withdrawal of the drug. Amphetamine if it may be required to overcome drowsiness.
2. Reserpine N.N.D. (Reserpan® Serp 12). Effective in even ly diabetics.
 - a. Give 2-5-5 mg ($\frac{1}{4}$ - $\frac{1}{2}$ gr) I.M. and tartaric acid solution 1 mg ($\frac{1}{80}$ gr) bid. Injection may have to be repeated daily every third day for 1-2 weeks. Maintain dosage of 1-2 mg daily thereafter.

- b Toxic reactions include moderate to severe depression (usually reversible on withdrawal of the drug) nasal congestion agranulocytosis and lethargy
- 3 Tranquilizing drugs recently introduced include
- Meprobarbital N N D (Miltown® Equanil®) For oral use only average dose 0.4-0.8 Gm (6-12 gr) q i d
 - Promazine Hydrochloride N N D (Sparine®) Suitable for intramuscular or intravenous use initial dose 50-150 mg ($\frac{3}{4}$ -2 $\frac{1}{4}$ gr) depending on degree of excitement. Thereafter intramuscular or oral doses of 25-100 mg ($\frac{1}{4}$ -1 $\frac{1}{4}$ gr) q i d
 - Prochlorperazine N N D (Compazine®) For oral use in doses of 5-10 mg ($\frac{1}{12}$ - $\frac{1}{6}$ gr) t i d
 - Perphenazine (Trilafon®) For oral use starting with 4 mg ($\frac{1}{30}$ - $\frac{1}{15}$ gr) t i d. Average daily dose 32 mg ($\frac{1}{2}$ gr). With higher doses extrapyramidal symptoms may occur.
- 4 Paraldehyde U S P is useful in delirium. Barbiturates, bromides and opiates often increase the excitement of delirium but may be used in maniacal states (see below). The ordinary stock paraldehyde solution needs no sterilization and for that reason is available for immediate administration by any desired route. The oral route is preferred unless the patient is unable to swallow. For details of administration see page 40.
- 5 Chloral Hydrate U S P may be given instead of paraldehyde in dose of 2-8 cc ($\frac{1}{2}$ -2 dr) of the 25% stock solution or as capsules 0.5-2 Gm (7 $\frac{1}{2}$ -30 gr) orally. It is contraindicated in acute alcoholic delirium or psychosis.
- 6 Barbiturates *not to be used for delirium*. Observe for respiratory depression and maintain adequate airway.
- Thiopental Sodium U S P (Pentothal Sodium®) First inject 3 cc of a freshly prepared 5% solution slowly I V observe then give additional dosage as needed for desired effect.
 - Amobarbital Sodium U S P (Amytal Sodium®) 0.125-0.5 Gm (2 $\frac{1}{2}$ -7 $\frac{1}{2}$ gr) as freshly prepared 10% solution slowly I V to point of desired effect.
- 7 Morphine Sulfate U S P 8-15 mg ($\frac{1}{8}$ - $\frac{1}{4}$ gr) with Scopolamine Hydrobromide U S P 0.3-0.4 mg ($\frac{1}{210}$ - $\frac{1}{160}$ gr) may be administered but when delirium is marked or associated with trauma caused by pain.
- 8 Scopolamine Hydrobromide U S P For delirium *without pain* scopolamine 0.3-0.4 mg ($\frac{1}{210}$ - $\frac{1}{160}$ gr) b i d q i d may be valuable.

D Hydrotherapy

- A warm tub bath (92-97 F) or soiled neutral bath for half hour periods t i d or q i d may be tried on suitable patients. This may be of considerable value. This method should be tried prior to instituting other therapy when possible. If it is tolerated well and results are effective the patient may remain in the tub for hours. Hydrotherapy is not applicable for certain unmanageable patients for patients with infectious or fibrile disease or for patients with surgical drainage.

- 2 Wet pack. This efficient technique should be administered only by trained personnel. The patient requires constant supervision. The method is contraindicated in patient who are physiologically exhausted, refractory convulsions or who have significant cardiovascular disease. Vital signs must be observed at least at intervals of 15-20 minutes.
- F Nutrition and Hydration. Unless there is specific indication for hypohydration a normal state of hydration should be maintained. This is especially true in the presence of fever. Food in moderate amounts of liquid 1.2 liters (1.2 qt.) of 5-10% glucose solution containing 100 mg (1/4 g.) of Thiamine Hydrochloride U.S.P. Aneurine Hydrochloride B.P. 100 mg Nicotinamide U.S.P. should be given daily. Protein intake should be maintained. Small frequent feedings are beneficial.
- F Prophylactic Calcium. If ordinary measures as mentioned above do not suffice consider transfer to psychiatric hospital. Evaluate feasibility of effecting the transfer and if decided upon provide for adequate attendant.

HEAD INJURIES

Proper management of the patient with a head injury rests in great measure upon thorough and accurate diagnostic and treatment methods.

Diagnosis

Careful initial examination and close observation of the patient in the immediate post-traumatic period are essential.

A Signs and Symptoms

- 1 Altered mental state of consciousness is the immediate period after the head injury. A 10-15 minute interval followed by coma may indicate cerebral compression by subdural or epidural hemorrhage. If progressively deepening consciousness after a period of consciousness following the injury, exploration of the skull is indicated to rule out subdural or epidural hemorrhage.
- 2 Progressive focal signs may indicate a dural hemorrhage.
 - a Ipsilateral pupil usually dilated.
 - b Contralateral hemiparesis may occur rarely.
- 3 If the patient remains unconscious diagnosis of a progressive intracranial hemorrhage is often difficult.
 - a Vital signs (pulse, temperature, blood pressure) may change although these are not reliable signs.
 - b In a state of deepening or ununusually prolonged coma, a plain roentgenogram of the skull is indicated. Prolonged unconsciousness is believed to indicate damage to the brain tissue.

B Laboratory Findings

- 1 Lumbar puncture is advisable to establish the presence of subarachnoid hemorrhage and to give baseline appearance and pressure of the cerebrospinal fluid.
- 2 Skull x-rays should be taken as soon as the patient's condition permits.

- a Presence and nature of fracture may be described
- b Presence of pineal shift can be ascertained
- 3 Electroencephalography may assist diagnosis and prognosis in selected cases in the chronic phase. Cerebral angiography may help demonstrate subdural or intracerebral hematoma

Treatment

A. Emergency Measures

- 1 Treat shock if present parenterally administered fluids and/or blood may be required (see page 29)
- 2 Attention to the respiratory tract is important maintenance of adequate airway and pulmonary ventilation is vital
 - a Patient should be placed in prone position with head turned to one side to facilitate drainage of secretions from mouth and to keep the tongue from obstructing the pharynx
 - b Intratracheal intubation or tracheostomy may be necessary to maintain open airway
 - c Give oxygen if necessary (see page 144)

B. General Measures

- 1 Quieting patient. During acute or initial phases restlessness may be a disturbing factor
 - a Special nursing care and paraldehyde may be required
 - b Avoid morphine because of medullary depressant effects
 - c Catheterization of a full bladder may melliorate restlessness
 - d Lumbar puncture with removal of small amount of bloody cerebrospinal fluid may also relieve agitated patient
- 2 Antibiotic treatment is always indicated in the presence of bleeding or discharge from nose or ears. Give Procaine Penicillin (U.S.P. 600,000 units) bid or broad spectrum antibiotic until discharge of infection is past

VASODEPRESSOR SYNCOPE

(Vasovagal Syncope Simple Fainting Benign Faint)

This is usually characterized by a sudden fall in blood pressure and a slowing of the heart. The causative stimuli may be sensory (e.g. sudden pain) or entirely emotional (e.g. death of a loved one). The patient is usually upright when the faint occurs and recovery rapidly restores consciousness.

Treatment

Patient should be placed in the recumbent position and head lowered. Simple inhalation of fumes of Aromatic Spirits of Ammonia (U.S.P. B.P.) may be tried if necessary.

ORTHOSTATIC HYPOTENSION

(Postural Hypotension) (code No. 460 x10)

This is a rare cause of syncope and occurs as the patient assumes an upright position. It is associated with a marked drop in blood pressure on arising.

Treatment

Treatment is directed towards the underlying cause when possible. If abdominal ptosis is present, an abdominal belt may prevent splanchnic pooling of blood. Elastic stockings may be of value. Vasoconstrictor drugs may be tried but are usually without benefit.

CAROTID SINUS SYNCOPE (code No 408 584 x)

There is usually a history of fainting associated with spells of distress between attacks. A definite relation to sudden turning or raising of head or wearing of a tight collar may be noted. The diagnosis is suggested by reproducing attack by firm pressure and massage over the carotid sinus for 10 to 20 seconds. Stimulate only one carotid sinus at a time. Caution must be exercised in stimulating the sinuses in elderly patients. Cerebrovascular accidents have been precipitated by this maneuver. Three types of carotid sinus syncope are known to occur.

Vagal Type

This is the most common type and is most frequent in older persons. Carotid sinus pressure slows the heart rate. This response can be abolished by the injection of Atropine Sulfate U.S.P. B.P. 1 mg (1/50 gr) I.V.

Vasomotor Type

Occurs more frequently in younger individuals. Carotid sinus pressure causes a fall in blood pressure. This can be abolished by injection of 0.5 cc (1/40 gr) of 1:1000 Epinephrine U.S.P. Adrenaline B.P. but is unaffected by atropine sulfate.

Cerebral Type

Carotid sinus pressure affects the heart rate and blood pressure and neither epinephrine nor atropine sulfate affects the reflex. A direct cerebral effect is postulated.

Treatment

Correct all abnormalities when ever possible. Eliminate emotional problems and forbid use of tight collars. In severe cases denervation of the sinuses may be necessary. Local anesthesia of the carotid sinus abolishes all type of carotid sinus syncope.

- A Vagal Type Atropine Sulfate U.S.P. B.P. 0.4 to 0.6 mg (1/160 to 1/100 gr) 3 or 4 times daily (or more frequently) will usually abolish attacks. Ephedrine Sulfate U.S.P. or Hydrochloride N.F. 25 mg (3/8 gr) with Phenobarbital U.S.P. 15 mg (1/4 gr) 3 or 4 times daily or Amphetamine Sulfate U.S.P. 5 to 10 mg (1/12 to 1/8 gr) may be used.
- B Vasomotor Type Ephedrine and phenobarbital is also effective usually prevents attack.
- C Cerebral Type Drug are of no value.

SYNCOPE DUE TO CARDIOVASCULAR DISORDERS

This type of syncope is due to cerebral anoxia, which results from a temporary fall in arterial output. Some of the causes are

Stokes Adams syndrome onset of paroxysmal tachycardia myocardial infarction and pulmonary embolism This may be associated with certain other types of heart disease (e g aortic stenosis and tetralogy of Fallot)

Treatment

Treat the underlying abnormality

SYNCOPE DUE TO METABOLIC DISTURBANCES

Hypoglycemia may cause syncope or coma If prolonged or recurrent treatment is required (see page 410)

Hyperventilation if severe and prolonged produces respiratory alkalosis with resulting tetany and syncope

Treatment

Consciousness can be restored by rebreathing into a paper bag holding breath or administration of carbon dioxide 5-10% with oxygen by mask If attacks are recurrent psychotherapy should be considered

VERTIGO

The term vertigo is generally used to denote the subjective sensation of rotatory movement either of the individual or his environment. Dizziness implies an inability to orient the body in relation to surrounding objects. However the terms are generally employed as synonyms. Vertigo is fundamentally in disassociation with involvement of the labyrinth the vestibular portion of the eighth cranial nerve and their neural connections. True vertigo is usually manifested by nystagmus falling to one side and abnormal reaction to tests of vestibular function. Among the more common causes are

1. Meniere's syndrome (see page 357)
2. Acute labyrinthitis (see page 357)
3. Organic brain damage involving the vestibular end organs or connections or the cerebellum
4. Drug and toxins (e g strychnine see page 507)

Treatment

Treat the underlying disorder

HEADACHE (code No 961)

Headache may be due to many factors and must always be recognized as a symptom. The underlying cause must be determined and treated in order to effectively relieve the symptom. The subjective sensation of headache indicates involvement of the pain sensitive structures within and about the skull. Headaches may be classified as follows with some of the more common causes listed

- A. Meningeal (and Allied Structures) Involvement. This is due to
 a. acute suppurative meningitis and a bacterial

p ssure a d d a d intracranial p ssure (following lumbar pun tu)

B V ul l vol em nt

1 l l r anial va dil tat on Due to feve c t in drug and toxins (e g al hol and hi tam) peis or motion l fa tor

2 Ext acranial va odilat t on (particularly f th xt renal carotid) M gr line is th p incipal example

3 Di eas of the blood v s els (e g t mporal arteriti)

C M s uloskeletal In ol m t

1 M s le ap m of v ryng degre s D e to myo ti ad j t thritis or o teiti

2 Muscle ten on du to emot onal f tor

3 B ne o joint in olvement of k ll h ad or cervical v rt b Due t arth itis o teitis o te my lit s or t mor

D Ne v l l m t N uralgias (e g t ig min l n uralgia)

E M ll o Ext ac l l n l m t Due t dise and disorde of th yes ears nose pharynx t th tc

F Emot on l l v lv m t H adache du t emotional d scord are u ally a t d with m sole te i n (e b ve) b t th s is t lways the case At times th h adache may be du t intracranial va d l t tio

D gno is

Th d gnosis must be ba d on a ompl te history and phy l l xaminat on Sp c al att ntion t the ey s and n is im portant A mpl t blood count urin lysis and blood test f yphilis must be pe f med In m ny case an ad qu te p y h at e amin t o s also und cated Skull y and l t o ncephalo g ms are u efu

Th pain f m ningeal involvem t i d ep a d is u ally th m st e P in f a ul igin is u ally throbbing in ch a ter pain of neuralgia has a burning qu lity He da h s of psy ho g i gin a p f ial and r man f st d by d ll tightness o p es u e

Ge l Non pe ifi T tm t M s

A Phy al and m t l st

B Sed tics should be used only as a t mp ary m ur and sh uld not be u d as a subst tute f a complete w k p and sp ifi th py Na c t are g lly ont und cated ex cept in terminal d s ase

C An lg i s on t but sp if th py m f b ll h d has due t thei antipyretic ctivity They should not be admini t d for p long dp od ind min t ly th ir r time u e sten ob cu s imp rtant p th l gy

HEADACHES DUE TO MENINGEAL INVOLVEMENT

Th e the m t e e e but th y usu lly e p d to an l g i Manif stat ona d p nd upon type and site f und lying p th lgy

T tm t

A Sp ifi M Tr t th th e lesion

B General Measures

- 1 Analgesics should be given as needed if pain is not too severe (see page 32)
- 2 Narcotics may be necessary if pain is very severe (see page 33)
- 3 Lumbar puncture performed very cautiously may sometimes be used to relieve headache associated with increased intracranial pressure (e.g. subarachnoid hemorrhage, hypertension, nephritis, not in posterior fossa tumors)

C Lumbar Puncture Headache These are believed to be due to leakage of the cerebrospinal fluid from the puncture site

- 1 Analgesics If headache is mild upon using analgesics such as Acetylsalicylic Acid U.S.P. (aspirin) 0.3 Gm (5 gr) q 3 hours may suffice. Codeine may be necessary
- 2 Recumbent position If lumbar puncture headache is very severe it can be alleviated by lying down
- 3 Intrathecal injection of small quantities of terbutaline may afford relief in severe cases

HEADACHES DUE TO VASCULAR INVOLVEMENT

These headaches are usually throbbing in character. Intracranial vasodilatation usually causes bilateral pain but migraine is usually unilateral. Compression of the common carotid may relieve both types of headache. Migraine may also be relieved by compression of the external carotid artery.

Treatment

A Intracranial Vasodilatation These are usually readily relieved by simple analgesics such as Acetylsalicylic Acid U.S.P. (aspirin) 0.3-0.6 Gm (5-10 gr) every 2-3 hours.

B Migraine (cod. No. 930 x40) Extracranial vasodilatation (be careful to avoid involvement of external carotid or its branches)

1 Treatment of an attack

a Ergotamine Tartrate U.S.P. B.P.

(1) Ergotamine tartrate 1 M strength and route of choice 0.25-0.50 mg ($\frac{1}{40}$ - $\frac{1}{20}$ gr) will relieve headache within an hour in most cases. Administer drug as early in attack as possible. Do not repeat dose more often than once weekly.

(2) Ergotamine tartrate by mouth 4 to 5 mg ($\frac{1}{15}$ - $\frac{1}{12}$ gr) sublingually or orally continued with 2 mg ($\frac{1}{30}$ gr) every hour until headache has disappeared or until a total of 11 mg ($\frac{1}{6}$ gr) has been administered. This method of administration is not generally advised because of the possibility of overdosage. If the patient vomits as a result of his disease it is impossible to know how much of the drug he has absorbed. Ergotamine is also less effective by the subcutaneous route.

(3) Toxic effects A few patients complain of numbness and tingling of extremities and some muscle pains and tension. Do not administer ergotamine to patients in septic or infectious states or who have peripheral vascular disease, arteriosclerosis, heart disease or who are pregnant.

- b Dihydroergotamine (D H E 45[®]) (not ac pt d) in dose of 1.0 mg (460 gr) i M or i V m y be substituted for ergotamine tartrate Injection may be repeated once h w r if necessary
- c Ergotamine with caffeine (Cafergot[®]) or atropine sometimes more effective by the oral route alone and requires a smaller total dose. It is valuable as suppo to less frequent use if emesis prevents oral administration
- d Pressure on external carotid or one of the branches early in the attack will often abolish pain
- 100% oxygen by nasal mask may relieve the acute attack
- 2 General measure
 - a Until drug begins to relieve headache have patient sit in a chair
 - b After headache has been relieved patient should rest in bed for the last two hours in a quiet darkened room without food or drink. This will promote relaxation and increase the possibility to prevent another attack from occurring immediately
- 3 Aborting an attack Many patients may abort their attacks by the following means
 - a Attempt to gain maximum relaxation by a warm bath
 - b Retire to bed in darkened room for several hours
 - c Drug
 - (1) Pentobarbital Sodium U S P Pentobarbital Sodium B P 0.1 Gm (1 1/2 g) by mouth
 - (2) Ergotamine Tartrate U S F B P 3.4 mg (1/20 g) bu g lly
 - (3) Acetyl Salicylic Acid U S P B P (aspirin) 0.6 Gm (10 g) with or without codeine 0.05 Gm (1 g) by mouth may be useful in mild attacks
- 4 Prevention of further attacks Migraine as a psychosomatic disease Reduction of attacks may be accomplished by psychotherapy There is little evidence that aspirin, diet, glandular therapy, antiallergic chemicals, etc. are effective except as psychotherapy devices
- C Histamine Antagonists (Hortons & Chahaligis) Since the use of 1.0 mg (460 g) histamine base (in histamine diphosphate solution) may sometimes reproduce the headache within 5 minutes Desensitization to histamine therefore has been demonstrated starting with a small dose of histamine diphosphate (0.25 cc) b i d and increasing the dose by 0.05 cc until a local edema is evident. Thereafter a maintenance dose of 1.0 cc 3 times weekly is injected
- D Diet of Blood Vessels Since many of the conditions are associated with a peripheral phenomenon vasodilators are indicated. Nicotinic Acid U S P 100 mg (1 1/2 g) i d t q i d or lly has been found to be of limited value

HEADACHES DUE TO MUSCULOSKELETAL INVOLVEMENT

Muscle contraction or spasms may be caused by disease of the muscle, adjacent structures or may be associated with cerebral fatigue or motion sickness. The muscle attachment to the occipital bone frequently involved gives the characteristic splitting

headache. There may also be a feeling of pressure or tightness or a band like constriction of the head associated with emotional tension. The psychogenic headache usually appears after periods of emotional stress.

Treatment

A. Muscle spasm due to organic disease and bone or joint pain may be relieved by appropriate physical therapeutic measures (see page 323). Analgesics are usually also of value (see page 3). Specific therapy should be directed at the underlying disease.

B. Muscle Tension Headaches

1. Rest, relaxation and freedom from emotional stress are of primary importance (see Psychotherapy on page 36).
2. Heat to the involved muscles by means of hot towels, heat ing pad or a warm bath will help relieve the discomfort.
3. Gentle massage of the muscles will usually also be of benefit.
4. Drugs may be of value in acute cases but prolonged use should be avoided.

- a. Phenobarbital U.S.P. Phenobarbitone B.P. 15
30 mg (1/4 1/4 gr) q. d. will temporarily relieve
many headaches due to nervous tension.
- b. Acetylsalicylic Acid U.S.P. (aspirin) 0.3 to 0.8 Gm (5
10 gr) every 3 to 4 hours may also be of benefit.
- c. Tranquilizers (see page 337 and 338).

THE DEGENERATIVE DISEASES

MULTIPLE SCLEROSIS (code No. 906.953)

A disease of unknown etiology characterized by patchy demyelination in the central nervous system which may be due to association with diffuse vascular thrombosis. It is manifested by diffuse neurological disturbances: cerebellar ataxia, nystagmus, slurred speech, intention tremor and spastic paralysis. C.S.F. examination shows nothing characteristic. The disease is slowly progressive with spontaneous temporary remissions.

Treatment

Chiefly symptomatic. Vasodilators (inhalation of 5-10% carbon dioxide, histamine infusions, amyl nitrite inhalations) are advocated for treatment of acute relapses by some experts.

A. Rest. Adequate sleep at night and rest in the afternoon has been found to make patients more comfortable.

B. Temperature Changes. Avoid sudden changes in temperature (extremes or internal) to reduce vasospastic phenomena (although evidence that pain plays a role in the disease is questioned by some). Heat makes the spasticism much worse, cold often improves them temporarily.

C. Rehabilitation. Physical therapy and psychotherapy to attempt to make the patient try to live with his disability and try to make the most of whatever assets he still retains.

PARALYSIS AGITANS (Parkinsonism) (code No 946 4 953)

A syndrom char act riz d by rhythmi al pill rolling tremor f sti g muscles with asso iated pa t city and igidity a tooped postu ma k i k fac s and a propulsive gait In later lif it is us ally as ocia t d w th art rioscleroti changes in the b sal ganglia In younge life it is usually n with po te ceph lit c chang s in the basal ganglia

T m t

Tr tment s m inly symptomatic Little c n be done to arrest the pr gres ive posten phalit c or arte ios lerotic hanges that o r

A Sp ific M a u e A numbe of drugs ha b n found to b eff ti in all viating the symptoms of parki sonism Th se drugs are usually used in ombinat on to obtain the optim l the ap tic result

C ti In patie ts with pa alysis agitans n ver stop one drug ab ptly when instit ting therapy with a new on Always int o du e th n w drug in sl wly in r sing qu t ti s whil grad u lly red ing the ld

- 1 Trih xyphe dyl Hyd o hlorid N N D (A tan ®) Eff c ti for stain d c trol of igidity mino tremor and akines Dos g 1 o 2 mg (1/60 1/30 g) to 5 mg (1/12 g) t i d Po oculogyric isis u 10 mg (1/6 gr) t i d Pri ipal ide act n same s for atropine but c do a ula ffe ts mi lm l Use w th ation i gla oma in high do age Artan ® m y cau e onf s o rest l ss e or hallu n tio s
- 2 Bell d nn ikaloids
At pine Solution of U S P B P (1/2%) Eff ct e for sp sma d gidity pa ticularly in post ncephalit s St t with 3 d p do s ti terval f about 6 h ur in r as dos g by l d pe y 3 days until a dos ge f 10 drops t d s e hed Limited to y ng r p tients be au of d ger of gl u om lde ly Ea ly t xic sympt ms blurring of vi ion dryn ss f m th ve tigo and t hy a dia Ec ive dosag m y p d m ting d un s me tal c nf si nd hallu i ations
- b Belladonna Tin t re U S P B P H s am ff t as atropin Start with 15 drop t d and in ease gradually t 30 d ps t i d Chief ld ffe ts a same s f at opin Do ot gl t pat e ts w th glaucoma
- 3 D ph hyd mi Hyd hio id U S P (B n d yl®) F t l of t mor Dosag 50 mg (3/4 gr) b i d to q i d
- 4 B n t pi M than llo t N N D (Cog ni n®) M t ff t e ge t g st igidity and p m lso againt t m E ll t wh e mbi d with t ih xyphe dyl (A tan ®) w ll as cy lmi (P gltan ®) o d tr mph t mi (De d ®) Chief sid tio dryne a f m th D ag B gun with 0 5 mg (1/120 g) 1 2 t mes daily and 1 cr d d p to 5 mg (1/12 g) daily
- 5 D t Amph t min S lfate U S P (De ed i ®) 5 mg (1/12 g) morning noo o Amph tamin S lf t U S P (B d i ®) 10 mg (1/6 g) to ou ter ct f tigue om n l e and l th gy

ANTISPASMODIC DRUGS

Drug	Effect On			
	Tremor	Rigidity (Sp. m.)	Alk. (w. k.)	Ocular Crisp
Atropine and belladonna alkali		x		
Belladonna (Cognate)	x	x		
Comphen (Proprietary)		x		
Diphenhydramine (Diphenhydramine)			x	
Diphenhydramine (Diphenhydramine)	x			
Ethopropazine (Proprietary)	x	x		
Rabellon	x			x
Rabellon (Proprietary)		x	x	
Serpa	x			
Trihexyphenidyl (Artane)		x	x	x
Mildred M. L. J. D. H. C. L. App. in L. F. Kin. T. m.				
Th. M. K. R. P. A. 1954 R. P. d. d. w. h. p. m.				

6 Hyoscine Hydrobromide U.S.P. B.P. Useful in control of tremor. Dosage ranges from 0.3 mg (1/200 gr) b.i.d. to q.d. in elderly patients to 0.6 mg (1/100 gr) b.i.d. t.i.d. in the young. Distressing side effects may include somnolence, dry mouth, blurred vision, and drowsiness.

7 Cycloman Hydrochloride N.N.D.† (Proprietary) Action similar to that of trihexyphenidyl (Artane) but has less drying effect. Useful where effect from trihexyphenidyl wears off. Dosage 12.5-50 mg (1/50-1/12 gr) t.d. to q.i.d.

8 Caramephen hydrochloride (Proprietary) Useful in young patients as muscle relaxant. Adult dosage 50-100 mg (3/4-1 1/2 gr) q.d.

9 Ethopropazine Hydrochloride N.N.D. (Proprietary) Lyko van 25-30 mg (3/8-3/4 gr) q.d.

10 Rabellon† (hyoscine hydrobromide atropine sulfate and acetylmethadol hydrobromide) Relatively little effect on tremor. Takes out 0.5 mg (1/120 gr) of mixed belladonna alkaloid. Give in 1/4-1/2 or full therapeutic dosage b.i.d. to q.i.d. depending on age and tolerance of patient. Side reactions: dryness of mouth and blurred vision.

11 Procyclidine Hydrochloride N.N.D. (K. Mader) Effective against rigidity. Does not produce the pronounced mouth dryness and blurred vision of some of the other drugs. Dosage 2.5-5 mg (1/24-1/12 gr) t.i.d.

12 Stramonium Tincture N.F. B.P. Especially good for control of tremor, tension, and excitement. Start with 15 drops t.i.d. and increase slowly to about 80 drops t.i.d.

13 Tranquilizers If patient is tense and anxious give Chlorthalidone Hydrochloride U.S.P. (Thalidone) 25-50 mg (3/8-3/4 gr) at bedtime. Reserpine N.N.D. (Reserpinol) Serpasil 0.25 mg (1/240 gr) t.i.d. or q.i.d. or Miltow b.m. N.N.D. (Miltow, Equil) 400 mg (8 gr) q.i.d.

†Contraindicated in glaucoma.

General Measures

- 1 Physical therapy Should include massage stretching of muscles and active exercises when possible. Patient should be taught to exercise daily the muscles most severely affected especially those of hands fingers wrists elbows knees and neck
- 2 Reassurance of control of symptoms and psychological support will be greatly helped by patient
- 3 Avoid harmful habits. Prohibit moderate use of alcohol some time for laxation

Prognosis

The disease is slowly progressive but it is not fatal

CEREBRAL VASCULAR ACCIDENTS

Cerebral vascular accidents are due either to thrombosis hemorrhage or embolism. The differential diagnosis is important in order to treat the underlying cause (see table below)

Differential Diagnoses of Cerebral Vascular Accidents

	Hemorrhage (34x 345)	Thrombosis (34x 318)	Embolism (34x 318)
Age	45-65 yrs	Over 45 yrs	Any age
Underlying cause	Hypertension	Arteriosclerosis	Cardiac diseases
Onset	Sudden	Sudden or progressively	Sudden
Headache	Absent	Slight or absent	Variable
Mental status	Coma	Normal to depressed	Normal to depressed
Paralysis	Complete hemiplegia	Slight partial hemiplegia	Slight partial hemiplegia
Spinal fluid pressure	High	Normal or slightly elevated	Variable
Blood in spinal fluid	Usually present	Usually absent	Absent or slight

Treatment

A. Acute Phase or Onset

- 1 Complete bed rest
- 2 Nursing care. Handle patient as fully as possible to avoid injury to patient and paralyzed extremities
- 3 Sedation (Paraldehyde USP) If patient is agitated sedation is necessary. However patient with thrombosis should not be depressed too much with sedatives
 - a Oral paraldehyde 1 (1 dr) in milk, fruit juice or whiskey repeated as necessary
 - b Rectal paraldehyde 8-15 (2-4 dr) in 30 cc of oil
 - c Intramuscular paraldehyde 8-15 (1-2 dr) deep into the buttock
- 4 Feeding. If patient is unconscious or unable to swallow do not attempt to give things by mouth. Maintain nutrition with tube feeding by parenteral means

- 5 Phlebotomy If hemorrhage has occurred and blood pressure is elevated phlebotomy of 500 cc may be used to reduce chances of further bleeding
 - 6 Lumbar puncture If hemorrhage has occurred perform lumbar puncture very cautiously removing just enough fluid to relieve severe headache Do not perform Queckenstedt's test in patients with suspected hemorrhage
 - 7 Voiding Catheterization may be necessary if spontaneous voiding does not occur
 - 8 Procaine block of the stellate ganglion has been recommended for thrombosis and cerebral embolism but is contraindicated in cases of hemorrhage
 - 9 Maintenance on anticoagulant therapy (page 215) which has been advocated for treatment and prevention of recurrences of cerebral thrombosis or embolism may be of value in thrombosis or insufficiency of the carotid or vertebral basilar system and cerebral embolism
- B State of Recovery and Convalescence** The rehabilitation of the patient with hemiplegia due to cerebral vascular accident should begin early and should be intense The details of the rehabilitation program are discussed on pages 327-334

Prognosis

If the patient survives the acute attack the prognosis for life may be good With active rehabilitation most patients will be able to walk and care for themselves Return of useful function to the upper extremity is rare (These patients can be trained to achieve a remarkable degree of recovery if given adequate care and rehabilitation) Prognosis for functional recovery is poor in those patients with severe residual organic mental syndrome or severe aphasia

HEPATO-LENTICULAR DEGENERATION (Wilson's Disease)

This extrapyramidal disease is characterized by progressive intention tremor athetosis rigidity dysphagia contractures muscle weakness and mental changes Flushing hemolysis and associated liver disease have been reported to be due to a defect of copper metabolism

Treatment

Dimercaprol U.S.P. B.P. (BAL®) has been reported to be effective in removing the excess copper The clinically effective dose is 2.5 mg (1/24 gr)/Kg body weight by injection b.i.d. for 10 to 12 days per course every 3 to 6 months

THE CONVULSIVE DISORDERS

EPILEPSY (Idiopathic) (code No. 934)

Epilepsy is a symptom complex which may be characterized by one or more of the following manifestations (Lennox)

- 1 Impairment of consciousness
- 2 Involuntary movement of muscles
- 3 Disturbance of the autonomic nervous system

Diagnosis

There are three major clinical types. The differential diagnosis is very important because the therapy of each differs. Individuals may have more than one type of seizure. Electroencephalographic study is indicated in all epileptic patients.

- A Grand Mal (code No. 930 x01) (Rule out the cause of convulsions.) This type occurs in all age groups. The usual form has generalized tonic and clonic convulsions which may begin focally and remain so or may spread without loss of consciousness (Jacksonian). The symptoms occur in single attacks varying in occurrence from hours to years.
- B Partial (code No. 930 x07) The usual form is characterized by a transient loss of consciousness of 5 to 30 seconds and generally no convulsive seizures. During the attack there is usually a rhythmic clasp and blinking of the eyes. It occurs most frequently in children and is rare after age 30.
- C Psychomotor Seizures (Epileptic equivalents) (code No. 930 x08) These forms frequently occur in adults and may be characterized by periods of abnormal behavior. The patient's emotional content is usually not altered from normal during the attack. The attacks vary in character and the patients are often dangerous to themselves and society.
- D Status Epilepticus (code No. 930 x06) Repetition of seizures of grand mal type which exhaust patient and may be fatal.

Treatment

Excitability epileptics not attempt given during an attack except to keep patient from being injured (e.g. biting his tongue).

A Grand Mal Never withdraw an anticonvulsant drug suddenly.

1. Diphenylhydantoin Sodium U.S.P. Phenylin Sodium B.P. (Dilantin®) is the drug of choice. Give 0.1 Gm (1½ gr) after evening meal 3 to 7 days in increasing degree by 0.1 Gm (1½ gr) daily every week until seizures are brought under control. If attacks are severe and frequent may begin with 0.3 Gm (4½ gr) daily on first visit. Average dose 0.4 to 0.6 Gm (6 to 9 gr) per day. After convulsions cease a patient treated with Dilantin® may be reduced if desired but avoid symptoms again as possible the dosage should immediately be raised again.
2. There are a few contraindications to Dilantin® but most troublesome is gum hypertrophy. This is best controlled with a careful mouth hygiene and gum massage. When large doses are given toxicodermatitis may appear (see p. 354).
3. Phenobarbital U.S.P. Phenobarbital B.P. If patient on maximum dosage of Dilantin® and there is inadequate response give phenobarbital in addition in same manner and dosage as Dilantin® in increasing dosage as with Dilantin® while maintaining patient at full dosage of Dilantin®.
4. Methyphenylhydantoin (Mephentoin®) If excessive gum hypertrophy results from the use of Dilantin® methyphenylhydantoin may be tried in its place. The dosage is the same. Mephentoin® may be effective where grand mal and

petit mal coexist. Do not suddenly change to Mesantoin® but gradually substitute for Dilantin®. Combination of both may prove more useful than the individual drugs. When using Mesantoin® special precautions should be observed for toxicity (see page 354).

- 4 Bromides: primidone (Mysoline®), mephobarbital (Mebaral®), benzchlorpropamide (Hibicon®) or ethosuximide (Peganone®) may be tried (see page 355).

B Petit Mal

- 1 Very mild state. If attacks are infrequent (less than 1 p day) give no treatment or treat only with small doses of phenobarbital.
- 2 Mild state.
 - a Amphetamine Sulfate U.S.P. (Benzedrine®) 5-10 mg ($\frac{1}{12}$ - $\frac{1}{6}$ gr) 2-3 times daily may be tried. Do not use if patient also suffers with grand mal because this drug may precipitate g and mal attacks.
 - b Glutamic acid 8-10 Gm (2 $\frac{1}{2}$ - 3 gr) daily may decrease the number of attacks.

3 Moderate and severe states

- a Trimethadione U.S.P. (Tridione®) is the drug of choice. Tridione is very effective in petit mal epilepsy but unfortunately is not an entirely safe drug since it causes bone marrow depression in some individuals. Whenever this drug is used perform CBC once or twice a week for the first month then every two weeks for two or three months and monthly thereafter. Dosage: Begin with 0.3 Gm (5 gr) daily and increase the daily dose by 0.3 Gm (5 gr) every 7 days until attacks are controlled. Do not give more than 2 Gm (30 gr) daily.
- b If g and mal seizures occur also trimethadione may aggravate this tendency therefore it may be necessary to administer medication for g and mal seizures simultaneously and in some cases atop the trimethadione.
- c Paramethadione (Paramadione®) is said to be less toxic than trimethadione. It is almost equally effective in petit mal attack and may be effective where other drugs fail. Observe precautions as for trimethadione (see page 354).
- d Phensuximide (Milonit®), phenobarbital (Mebaral®), ethosuximide (Dilantin®), acetazolamide (Diamox®), mephobarbital (Mebaral®) may prove useful (see page 355).

C Psychomotor Epilepsy Patients must be watched and guided to prevent injury to themselves or others.

- 1 Diphenylhydantoin Sodium U.S.P. Phenytoin Sodium B.P. (Dilantin®) without phenobarbital as first and main epilepsy is the treatment of choice.
- 2 Phenacemide N.N.D. (phenylacetylurea Phenazone®) is effective in control of psychomotor epilepsy. Give initially 0.5 Gm (7 $\frac{1}{2}$ gr) t.i.d. and increase until symptoms are controlled up to 5 Gm (75 gr) daily divided into 3-5 equal doses. The drug is quite toxic and precautions must be observed with its use (see page 354).
- 3 Methyphenylacetate (Mesantoin®), mephobarbital (Mebaral®), primidone (Mysoline®), acetazolamide (Diamox®)

a d methsuxim d (C lout n®) alone or in combin tion with
oth dugs a f que tly seful

D Stat Epilepti

- 1 Am ba b i l S d um U S P (Amytal Sod m®) 0 5 1 Gm
(7½ 15 g) I V may be g v I t ave ous Ph b b
ial Sod m U S P Ph n b r b t Sodium B P 0 4
0 8 Gm (6 12 g) in j i d l wly m y b e u s d
- 2 Paral d hyd U S P 1 2 cc d l t e d in t p l e volume of
aline I V lowly If onvulsion ontin e e p e t l V dos
t t V E R Y S L O W L Y A N D C A U T I O U S L Y o r g v e 8 12 I M
- 3 D p h e y l h y d a n t o l S d m U S P Ph y t o l n Sod m B P
(D i a l t i n Sod m®) may b i n j e c t d i V a t a t e n o t e d
g 50 mg (¾ gr) p e r m u t e A t o t a l d o g e o f 150 250
mg (2½ 4 g) may b r e q u i r e d
- 4 G n e r l a n e s t h i a m a y b e e d f a l l o t h e r m e a s u r e f a i l
- 5 D i a l t i n® A s s o o e d t v m e a s i f c t i v p a s s
t o m a c h t b e t h r o g h n o e e n i f p t i e t s t i l l s c i o u
a d g 0 1 0 2 Gm (1½ 3 g) D i t i n® i w a t e r y
3 h o u r s t i l s e i z c o t l l e d (m a x m u m 10 d o s)

E D t o n o f T r e t m t M o t p i l p t i m t i e t h a p y
f l i f H o w e e i f i s r t r l y o t l l e d f o 3 5
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F G r l M e s e s A c q u t t h e p t i t w i t h h d s I n
s t b l e c a s t h m y b a c o m p l e d n p t b y r a d g
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p o t a n t t o m a i n a r e g u l p o g r a m f p h y i c a l t i v i t y
K p t h e p t t i o p t m m p h y l c o d t i o d a v o i d e x
e s s f t g e F b d a l l a l h o l T a t e m t i o a l f c t o
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H p a d B s { } P t m O n C o n v u l s v e S u e s M
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Identificati n Card

THIS PATIENT HAS EPILEPSY

Name _____
Add s _____ Phon _____
P t a p a d d t k o p a d d p o o i n h s m o u t h t o
p t t h t g u K p h m f r o m i n j r g h i m s I f
D t s N m e _____
Add s _____ Phon _____

Ph ob b i	All p i p	0 1 0 + Gm	1 D in	1 D d	On i ad d g M y
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		di id d d		top d g	
Ph m d	P t t m i	0 6 2 4 Gm	1 N	1 R d d g	
N N D			2 At i	2 d i onv	
(Nation y)			3 D i in	4 D s t i	
U h m d	P m a l	1 2 Gm	1 At	dos	
(C i y)	P y h m i	(0 3 Gm	2 D w i		
	p i p	9 i d)	3 C a i h	2 d i t i	
A i m d	G d m a l	1 3 Gm (0 25	1 D	R d i	
N N D (D am y)	P i m i	Gm i d)	2 P t h i		
E h in N N D	G d m a l	2 3 Gm 4 d	1 D a	1 R d d	
(P g a n o n y)		d d d d	2 P i g	2 d i i	
		f m i	3 S k i h	3 d i i	

NARCOLEPSY

Narcolepsy is a clinical syndrome characterized by recurrent episodes of uncontrollable desire to sleep. It is frequently associated with a transient loss of muscle tone (cataplexy) especially during emotional reactions.

Treatment

- A Amphetamine Sulfate U.S.P. (Benzedin®) 10-175 mg ($\frac{1}{8}$ to $2\frac{3}{4}$ gr) daily may be required. Optimum dosage may be determined by starting with dosage of 10 mg ($\frac{1}{8}$ gr) each morning and increasing by 10 mg increments through the day. 15 to 20 mg ($\frac{1}{8}$ to $\frac{1}{2}$ gr) t.i.d. is an average dose.
- B Ephedrine Sulfate U.S.P. Ephedrine is not as satisfactory as amphetamine but is helpful in many cases. Optimum size and distribution of dose varies with patients. Average range of dose is 25-50 mg ($\frac{3}{8}$ to $\frac{3}{4}$ gr) b.i.d. q.i.d.

DISEASES OF THE CRANIAL NERVES

TRIGEMINAL NEURALGIA (code No. 954 x30)

Trigeminal neuralgia is characterized by sudden attacks of excruciating pain of short duration anywhere along the distribution of the 5th cranial nerve. The attacks are normally precipitated by stimulation (usually mild) of the trigger zone with each of the pain.

Treatment

- A Medication treatment is generally unsatisfactory but the following usually are tried before resorting to surgery.
- 1 Trichloroethyl ether U.S.P. (Trilene®) 15-20 drops per day by inhalation from the dark blue funnel or directly into the nose one-half hour before meals.
 - 2 Magnesium sulfate B.P. (2000 mg magnesium daily by injection for 10 days) has been reported to relieve the pain of trigeminal neuralgia.
 - 3 Sodium diethylenetriamine N.N.D. has been shown to produce a chemically neuropathic effect on the facial and trigeminal ganglia. G 0.15 Gm ($\frac{1}{2}$ gr) freshly dissolved in 10 cc 5% glucose and distilled water i.v. over a period of one-half hour. Attacks of 10 daily injections recommended. Relief may be delayed for 5 months until the chemically neuropathic effect is well pronounced. Side effects include drowsiness, dizziness, and some of the following:
 - a Anticonvulsants e.g. Diphenylhydantoin Sodium U.S.P. (Dilantin®) 0.1 Gm ($\frac{1}{2}$ gr) q.i.d. or sodium valproate.
 - b Tolazemide Hydrochloride U.S.P. (Pavoline®) 50 mg ($\frac{3}{4}$ gr) q.i.d. have been reported to be helpful in some cases.
- B Surgery may be required if the above fails from medical treatment.

BELL'S PALSY (Peripheral Facial Paralysis) (code No 865 y10)

A palsy is of all the muscles of one side of the face usually precipitated by exposure to chill or to sun.

Treatment

After the patient has recovered usually occurs gradually in 2-8 weeks it may take up to 12 years in older patients.

A Protection of Face

1. Keep face warm and avoid further exposure
2. Protect eye with a patch if necessary
3. Avoid wind and dust

B Physiotherapy

1. Support face by use of tape or wire if necessary of mouth looped about the ear
2. Electrical stimulation may be used to help prevent atrophy of muscles. Use every 2 days after the 14th day
3. Gentle massage in an upward direction for 5-10 minutes 2-3 times daily of the involved muscles may help the tonus
4. Heat from infra-red lamp may help to relieve

MENIERE'S SYNDROME (code No x00)

Meniere's syndrome is a symptom complex of unknown etiology which involves the labyrinthine portion of the 8th cranial nerve. It is manifested by sudden recurrent attacks of vertigo, nausea, vomiting, nystagmus, and tinnitus and by progressive deafness.

Treatment

A Special Measures Non available

B General Measures

1. Rest is essential. Many of these patients have marked psychoneurosis.
2. Salt restriction and Ammonium Chloride U.S.P. B.P. 12 Gm (15-30 g) q.d. may be helpful.
3. Nitroglycerine U.S.P. (in oil tin) 50-100 mg ($\frac{3}{4}$ - $1\frac{1}{2}$ g) i.v. bid to tid 100 mg ($1\frac{1}{2}$ gr) orally 5-6 times daily has been found useful.
4. The antihistamines especially Diphenhydramine Hydrochloride U.S.P. (Benadryl®) and Dimenhydrinate U.S.P. (Dramamine®) in doses of 50-100 mg ($\frac{3}{4}$ - $1\frac{1}{2}$ gr) 3-4 times daily appear to be of benefit to some patients.
5. Section of the vestibular portion of the affected 8th cranial nerve may provide relief in cases not responsive to medical treatment.

DISORDERS OF EQUILIBRIUM

ACUTE LABYRINTHITIS (code No x85 910)

Acute labyrinthitis is an acute inflammation of the inner ear which usually follows purulent infection and is manifested by intense vertigo usually with marked tinnitus and staggering gait and nystagmus.

TreatmentA Specific Measures None availableB General Measures

- 1 Bed rest preferably in darkened room until symptoms subside
- 2 Drugs
 - a Antibiotics are of little value unless there is a local infection of middle ear or mastoid
 - b Antihistamine drugs may be of some value (as for motion sickness see below)
 - c Sedation is generally helpful Phenobarbital U.S.P. Phobarbitone B.P. 15-60 mg ($\frac{1}{4}$ to 1 gr) t.i.d. to q.i.d.
 - d Chlorpromazine Hydrochloride U.S.P. (Thorazine[®]) 50 mg ($\frac{3}{4}$ gr) i.m. (or other phenothiazine derivative see page 42) is useful in the acute early phase

MOTION SICKNESS (code No. 010 576)

Motion sickness is an acute illness characterized by anorexia, nausea, dizziness, and vomiting. Many factors play a role in its production; the principal ones being visual kinesthetic and psychological. Physiologically the vestibular apparatus appears to be involved.

Prophylaxis

Preventive measures are often effective. Attacks of motion sickness are difficult to treat successfully.

- A The antihistamines appear to be of benefit. Dimenhydrinate U.S.P. (Dramamine[®]) or Diphenhydramine Hydrochloride U.S.P. (Benadryl[®]) 50-100 mg ($\frac{3}{4}$ to 1½ gr) q.i.d. is said to be very effective.
- B Meclizine Hydrochloride N.N.D. (Bonine[®]) 25 mg (½ gr) tid effective against the usual dose is 50 mg ($\frac{3}{4}$ gr) tid 12 hours.
- C Cyclizine Hydrochloride N.N.D. (Marz[®]) is effective orally i.m. doses of 50 mg ($\frac{3}{4}$ gr) repeated 4-6 hours p.r.n.
- D Parasympathetic depressants also are in combination with mild sedatives. Scopolamine Hydrobromide U.S.P. or Atropine Sulfate U.S.P. 0.2-0.4 mg ($\frac{1}{300}$ to $\frac{1}{150}$ gr) tid 3-6 hours.
- E Mefenazine Hydrochloride U.S.P. Phobarbitone B.P. 15-30 mg ($\frac{1}{4}$ to $\frac{1}{2}$ gr) tid 3-6 hours may help prevent attacks.

PERIPHERAL NEURITIS (code No. 08 910)

Peripheral neuritis can be caused by a large number of factors, both local and general. There may be either a non-inflammatory (with pain, paresthesias, and therapeutic response to symptomatic treatment) or motor involvement (with weakness and paralysis) but motor frequently both.

- 1 Toxic Factors E.g. lead, arsenic, mercury, alcohol, etc.

- B if t g Goll in B e type of multiple neur t
 C Dilecy Typ Esp lly of th B compl (be b i) i oft
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Tr im t

I m nt in h d pends p n th etiologic l f tore

A Sp c f T m t

- 1 R mo nox ous agent g l chol l d sour
 V tam B compl Att mpt t obtai optim i met bol m
 f ner t e by l b alus f v t m espe lly B compl
 pl x Thiam e Hyd o hlo de U S P A i e Hyd
 hlo id B P 15 mg (1/4 g) t i d to b i d o lly o
 p te ally d D d Y st U S P (b w s y l)
 10 30 Gm (1/3 t) daly f e t i e B compl (se p ge
 62)
- 3 A t t h gh l r i c d t i i n d t d
- 4 I l d polyn ritis of ch i t g a g t such a Ed th
 m i C l m d dium N N D (EDTA V e ate[®]) m y
 b be f l (p g 54)
- 5 I e scal p lyneur t s Dum cap l U S P B P
 (BAL[®]) (se pag 536)

B G l T m t

- 1 B d t P l c pat t b d if po ibl a d avo du of
 ff t d l m b If low tr m ty aff t d k p adi
 ov fo t f bed to p e e t p e sur of bed c e
 A lgesics e e s y to c t l p (se p g 3)
- 3 Phys l th apy (page 333) Aft p has sub d d
 phy l th r py (m e s g a d p i m t i o) m y b e of
 v l e E ou ag t e moti n at th s m time P e t
 i t e s by m ans of plant d p at t h i g

HERNIATION OF INTERVERTEBRAL DISK (code No 2511 9x9)

Comp es and i jury t e oot may b ca s d by h ni
 t n f an nt rv t b r a l d k M t commonly th lumb s l
 i t e t b a l d k (L5 S1 L4 L5) s aff t d in th ca s s
 th ymptom ompl of low b k p un impa d a g f mot o of
 low r b ck p av teb l i m b a m u c l p sm and pain disting
 l g th iat d i t b t i n commonly ounte d O s t f
 l i n i c l ompl i t e s f qu t i y r l t d to p o d s m v l i n g low
 b k t i back injur of f l l Th initi l p i d m y b
 f l l w d by an t val f ympt mat improv m nt

Exagg at on of symptom t i c mplaunt f l l o w f qu t i y p o n
 la ghung i t a u n g s n z i n g

T d s i n s a t i c t h a l g e o u of th c t i n r
 m p i r d t a g h t i g r a i s i n g d m u n h e d n k l e j k i m p a d s n
 t o o d s t i b t i o n of L5 of S1 may be dem nstr t d

Ch a t i a t i o e t g l g l d f t i n th b m d p a c
 s lly p o d d b y h i d i n t e r v t b l d i s k a n d e d i l y
 d m o n t b l by m y l o g r p h y

TreatmentA Specific Measure None availableB General Measure

- 1 Bed rest preferably in darkened room until symptoms subside
- 2 Drugs
 - a Antibiotics are of little value unless there is associated infection of middle ear or mastoid
 - b Antihistamine drugs may be of some value (as for motion sickness see below)
 - c Sedation is generally helpful Phenobarbital U S P Phenobarbitone B P 15-80 mg (¼-1 gr) t.i.d. to q.i.d.
 - d Chlorpromazine Hydrochloride U S P (Thorazine®) 50 mg (¾ gr) i.b. (or other phenothiazine derivatives see page 42) is useful in the acute early phase

MOTION SICKNESS (code No 010 576)

Motion sickness is an acute illness characterized by an acute nausea, dizziness and vomiting. Many factors play a role in the production, the principal ones being visual kinesthetic and psychological. Physiologically the vestibular apparatus appears to be involved.

Prophylaxis

Preventive measures are often difficult. All kinds of motion sickness are difficult to treat successfully.

- A The antihistamines appear to be of benefit Dimhydrinat U S P (Dramamine®) or Diphenhydramine Hydrochloride U S P (Benadryl®) 50-100 mg (¾-1½ gr) q.i.d. said to be very effective.
- B Meclizine Hydrochloride N N D (Bonamine®) a long acting effective agent. Therapeutic dose: 50 mg (¾ gr) every 6-12 hours.
- C Cyclizine Hydrochloride N N D (Marion®) a highly effective oral dose of 50 mg (¾ gr) repeated in 4-6 hours as required.
- D Parasympathetic depressants also of combination with meclizine: Atropine Hydrobromide U S P or Atropine Sulfate U S P 0.2-0.4 mg (1/300-1/150 gr) every 3-6 hours.
- E Mild Sedation Phenobarbital U S P Phenobarbitone B P 15-30 mg (¼-½ gr) every 3-6 hours may help prevent attacks.

PERIPHERAL NEURITIS (code No 98 710)

Peripheral neuritis can be caused by a large number of factors, both local and general. There may be either sensory in character (with pain paresthesias and other subjective sensations) or motor involvement (weakness and paralysis) but more frequently both.

A Toxic Factors. E.g. lead, arsenic, mercury or diphtheria toxins.

therapeutic test of Neostigmine Methylsulfate U S P (Prostigmin[®]) 1.5 mg (1/45 gr) with Atropine Sulfate U S P 0.6 mg (1/100 gr) (diagnostic ampul) subcut may be used. This causes relief of symptoms.

Treatment

A Emergency Treatment Patients may suddenly develop inability to swallow or respiratory distress. Patient should always carry 2 ampuls of 0.5 mg (1/20 gr) of Neostigmine Methylsulfate U S P B P. This should be given immediately subcut or IM if severe symptoms develop. The patient should be placed under medical care at once and if additional neostigmine is needed 1 mg (1/80 gr) may be given parenterally 2-3 times in one hour until adequate response is obtained.

In spite of administration of increasingly large amounts of neostigmine, patient will have sufficient muscular respiration may occur which may be fatal in some cases.

1. When such an event can be anticipated, tracheotomy and oxygen equipment, suction apparatus and respirator should be available.
2. Following tracheotomy, patient is placed in respirator. Oxygen administered. If Neostigmine is withheld, Atropine Sulfate U S P 0.6 mg (1/100 gr) is given to keep airway dry. Intubation of airway employed.
3. Maintaining fluid and electrolyte balance during artificial respiration. Give antibiotic to prevent pneumonia.
4. After a few days, it is usually possible to gradually discontinue patient in the respirator as tolerated.
5. Patient who develops myasthenic crisis may occur in some instances, usually severe.

B Specifics

1. Neostigmine Bromide U S P B P 15 mg (1/4 gr) tablets. Dosage: 1 to 4 tablets daily. Begin with 1 tablet every 4 hours (4 times a day) and increase as required to give relief.
2. Ephedrine Sulfate U S P 12 mg (1/5 gr) with the help of neostigmine facilitates intubation of trachea.
3. Patient may have labored breathing due to pleuritic rigidity. It must however be given in close therapeutic doses 4-6 Gm (60-90 gr). Potassium Chloride U S P 8 times a day. This is indicated if digitalis is being used.
4. Edrophonium Chloride N N D (T Nallo[®]) may relieve myasthenic weakness. 10 mg I V gives relief in 30 seconds. 25-50 mg I M gives longer lasting relief for hours. Two to 3 mg (1/30-1/20 gr) I V may be used as a test dose for patient to determine time to distinguish between the effects (impulse) and orthostatic (change).
5. Pyridostigmine bromide (Mestinon[®]) a analog of neostigmine is at times more effective in the treatment of bulbar muscle weakness. It is applied in 60 mg (1 gr) tablets. Dose: 10-25 tablets daily. Intubation is preferred to produce maximal relief.
6. Ambesonium chloride (Mytilas[®]) is said to act twice as long as neostigmine and has fewer side effects. Start with 5 mg (1/12 gr) tid and increase as needed. Average dose 5-25 mg (1/12-3/8 gr) qid.

380 Myasthenia Gravis

Treatment

A General Measures

- 1 In acute phase bed rest heat applied locally to back and girdles and use of a bed board under mattress are indicated
- 2 Traction to the lower extremities is frequently beneficial
- 3 The avoidance of severe physical effort and strain is essential to minimize recurrence of symptoms after the initial episode
- 4 Use of low back belts braces or supports may be beneficial It is important to instruct patient as to proper method of bending lifting (with knees flexed) and carrying (with object held close to body)

B Surgical Measures Where response to conservative measures is poor or when recurrences have disabled the patient surgery is indicated Gratifying relief of the major complaint of most patients i.e. pain usually follows the successful removal of the offending herniated disk Recovery of other neural functions (impaired motor power muscle atrophy skin sensory changes) may be expected later

MYOPATHIES

MYOTONIA

(Congenital code No 270 044) (Acquired code No 270 x20)

Myotonia is a disorder characterized by difficulty in relaxation of skeletal muscles following contraction which is initiated either by voluntary effort or by mechanical or electrical stimulation It is important to differentiate this disease from myasthenia (see below) because treatment with neostigmine or potassium aggravates myotonia

Treatment

Quinidin Sulfate USP BP 0.306 Gm (59 gr) 3-4 times daily may be used to relieve symptoms

PROGRESSIVE MUSCULAR DYSTROPHY (code No 270 9x0)

A disorder characterized by progressive wasting and weakness of muscles with associated peripheral hypertrophy (fat infiltration) of certain muscle groups It is important to differentiate this from myasthenia gravis because the latter can be benefited by treatment

Treatment

None of value It has been suggested that inability to metabolize vitamin E may play a role in the disease but parenteral administration of this substance has been of no benefit

MYASTHENIA GRAVIS (code No 270 562)

Myasthenia gravis is a disorder characterized by weakness and marked fatigability of voluntary muscles Recovery from weakness or fatigue occurs within minutes or hours of rest The disease progresses by natural or spontaneous remission and relapses

Chapter 15

METABOLIC AND ENDOCRINE DISEASES

DISEASES OF THE PITUITARY

In the diagnosis and treatment of endocrine disorders it must be remembered that there is a very close interrelationship of the various endocrine glands. Not only do hormones exert profound effects on all tissues of the body but the endocrine glands also exert strong influences upon each other. For this reason the manifestation of endocrine diseases may be either primary to a given endocrine gland or secondary, i.e., due to involvement of a target gland. For example, the patient with hypopituitarism may present with a pituitary adenoma and unless it is appreciated that the primary disturbance is in the pituitary gland, the treatment of thyroid disease without the simultaneous use of corticosteroids is a serious disadvantage.

PANHYPOPITUITARISM (code No. 841 777) HYPOPITUITARY CACHEXIA (Simmonds Disease) (code No. 841 7773)

Organic hypopituitarism is due to destruction of the pituitary which may be caused by tumors of the gland or postpartum hemorrhage (Sheehan's syndrome).

The term panhypopituitarism is probably a misnomer since the significance of variations in symptomatology of the disease may be due to varying degrees of the several hypopituitarisms. Hypopituitary cachexia is also a misleading term since the patient may be of normal weight and may actually be obese. Symptoms of organic hypopituitarism usually include weight loss, weakness, sensitivity to cold, loss of appetite and infrequent menses or amenorrhea. Physical examination reveals the signs of axillary and pubic hair and atrophy of the kidneys and genitalia. Laboratory findings are slow basal metabolic rate, low basal iodine uptake, PBI is decreased, urinary gonadotropin is low, and urinary gonadotropin is low. These manifestations are due largely to lack of pituitary gonadotropin secretion. The treatment is with thyroid gland extract.

362 Familial Periodic Paralysis

- C General Treatment Acquaint patient with his disease using simple lay terms. Maintain good nutrition and health.
- D Surgery Thymectomy is claimed to benefit some patients.

Prophylaxis

Identi fication card, neostigmine and a syringe are to be carried at all times giving the diagnosis and method of treatment.

THIS PATIENT HAS MYASTHENIA GRAVIS

Name _____

Address _____ Phone _____

If observed to be behaving strangely or if he is found unconscious call for physician or ambulance.

Physician's Name _____

Physician's Address _____ Phone _____

Two ampule of neostigmine and a syringe and needle are in his possession. These must be administered hypodermically into his upper arm immediately.

Management During Pregnancy

Immediately after delivery children of patients with myasthenia may have severe signs of the disease. Immediate treatment with neostigmine is necessary to preserve life. After a few days the symptoms may disappear and the child thereafter does not suffer from myasthenia.

FAMILIAL PERIODIC PARALYSIS (code No. 270 x95)

A disease of unknown etiology characterized by recurrent attacks of flaccid paralysis of the muscles of the trunk and extremities and by a lowering of the serum potassium level during the attack. Immediate relief of symptoms by administration of potassium chloride is usually diagnostic.

Treatment

- A Potassium Chloride U.S.P. B.P. 5-10 Gm (75-150 gr) orally when diagnosis has been made and then 5 Gm (75 gr) b.i.d. q.i.d. during acute episode as needed to prevent weakness or paralysis.
- B In emergencies only may give prepared solution containing 1 Gm (15 gr) Potassium Chloride U.S.P. B.P. in 50-60 cc distilled water injected very slowly I.V. This is a dangerous procedure (see p. 25).

Prophylaxis

- A Avoid high carbohydrate foods (e.g. candy, honey, sugars).
- B Routine administration of 25% aqueous solution Potassium Chloride U.S.P. B.P. (or enteric coated tablets) 5-12 Gm t.i.d.

GIGANTISM (code No 841 7781)

Pituitary gigantism which is caused by adenoma of the anterior of the pituitary is principally a result of hypersecretion of the growth factor Growth which occurs primarily at the epiphyses of long bones is symmetrical and generalized and patients may attain a stature of several feet high Growth is possible only if the oversecretion of hormone occurs prior to the onset of epiphyseal closure. Lethargy is a symptom of the pituitary tumor may cause headache, hypertension, and visual disturbances. An elevated serum inorganic phosphorus is one of the best diagnostic signs of a pituitary Glycosuria may be present.

Treatment

- Treatment is aimed at suppressing the pituitary growth hormone.
- A. Endocrine Therapy If gigantism is found in adolescence and the patient is a male, the treatment of the tumor posteriorly is indicated. (1) Optimal dose is given at a single dose of 400 mg (420) 400 mg by mouth daily. Ethinyl Estradiol USP 0.1-0.5 mg by mouth daily. Males should be followed for the first 6 months of treatment. Females should be followed for the first 6 months of treatment. Do not use methyltestosterone.
- B. Surgery and X-ray Therapy If the tumor is not removed surgically, the patient should be followed for the first 6 months of treatment. If the tumor does not regress, growth surgery must be considered.

ACROMEGALY (code No 841 7782)

Hypersecretion of the growth factor of the anterior pituitary due to adenoma of the gland which develops after the bone epiphyses have fused results in clinical picture of progressive growth of soft tissues and thickening of bones. The disease usually has onset during the 2nd or 3rd decades. It is characterized by enlargement of the jaw bones, supraorbital ridges, hands and feet, thickening of the skin, and visual impairment. Elevation of the inorganic serum phosphorus is a important diagnostic point if the serum phosphorus is normal, the disease is probably inactive and requires no further treatment.

Treatment

Acromegaly (see above) Favorable results have been reported in some cases with endocrine therapy.

Differentiation of panhypopituitarism from anorexia nervosa (functional hypopituitarism) may be difficult. Psychic data base may be found in both conditions although a history of specific emotional stress or long standing psychiatric symptoms is more suggestive of anorexia nervosa. The nervosa patient is usually more alert and active and more able to withstand stress. The axillary hair is usually not lost in anorexia nervosa but is almost always lost in organic hypopituitarism. The low urinary gonadotropin level (less than 3 mouse units per 24 hrs) of hypopituitarism may be of aid in diagnosis but is not definitive. Improvement following special feeding technique and failure to respond to specific endocrine treatment would further suggest anorexia nervosa.

Treatment

There is no effective pituitary replacement preparation. Therapy must therefore be aimed at correcting the end organ deficiencies. This must be continued throughout life. Almost complete replacement therapy can be carried out with cortisone.

A Cortisone or Hydrocortisone 7.5 to 25 mg per day is usually adequate. This should be given in divided doses 3-4 times daily.

B Thyroid Thyroid (and insulin) should rarely if ever be used in panhypopituitarism unless the patient is receiving cortisone. Because of lack of adrenal cortical function patients are exceedingly sensitive to these drugs. For this reason one should exercise special care in differentiating myxedema from hypopituitarism, often a difficult problem.

Begin with small doses of 15 to 30 mg ($1/4$ to $1/2$ gr) daily and gradually increase to tolerance 60 to 100 mg (1 to $1\frac{1}{2}$ gr) is usually adequate.

C Sex Hormones

1 Testosterone May be used in both males and females primarily for its tissue building (protein anabolic) effect. Dosage One of the longer acting parenteral testosterone preparations (see page 40) 100-300 mg every 3-4 weeks. Methyltestosterone 15-30 mg B.P. 10 to 20 mg orally in males; a female with dose of these drugs is half that for males. If virilizing signs appear in the female the drug should be stopped and these signs will disappear. Virilizing signs usually do not result if the dose is kept under 400 mg per month.

2 Estrogens These agents are useful in the female for their mild anabolic effect, their effect on secondary sex characteristics and their possible neutralizing effect on androgens. Diethylstilbestrol U.S.P. 1 mg or Diethylstilbestrol B.P. 0.5 to 0.6 mg or Ethinyl Estradiol U.S.P. 0.02 to 0.5 mg daily or orally.

Note Sex hormones especially estrogens should be employed cautiously in young hypopituitary patients or the epiphyses will close before maximum growth is achieved.

probably necessary. At times of stress, especially in puberty and during pregnancy and lactation, the requirement rises a high as 200-1000 micrograms (0.2-1.0 mg) daily.

Abnormal Metabolism

Although the iodine requirements are very slight, in many areas of the United States and elsewhere, these requirements cannot be met from local food and water sources.

A Simple Iodine Lack (Simple Goiter) Endemic goiter or colloid goiter is characterized by enlargement of the thyroid gland and is due to relative or absolute iodine deficiency with a secondary work hypertrophy of the gland. There is often a history of living in an iodine deficient geographic area. Symptoms appear only if the enlargement is sufficient to cause pressure on surrounding structures (esophagus, trachea, or recurrent laryngeal nerve). The incidence of either hyper- or hypofunction and accordingly the BMR, serum protein-bound iodine and cholesterol and diiodine (I^{131}) uptake are usually normal.

B Hypothyroidism In this condition the gland fails to manufacture adequate hormone. This may have various causes: (1) mere complete iodine lack, (2) inflammatory destruction of the gland (thyroiditis), (3) excessive surgical removal, (4) failure of the pituitary to elaborate thyrotropin. In hypothyroidism the BMR, radiiodine (I^{131}) uptake and blood organic iodine are frequently low (the latter below 4 micrograms per cent).

C Hyperthyroidism This disease is characterized by an excessive secretion of thyroid hormone. The causes of this are obscure, but it is believed that in many cases the primary difficulty may be excessive secretion of anterior pituitary thyrotrophic hormone. This excessive secretion causes a speeding up of metabolic function, especially the oxidative mechanism of cells. This is a result of an excess of BMR, the blood levels of organic iodine are frequently above 8 micrograms per cent, and the I^{131} uptake is high.

DISEASES OF THE THYROID

NON TOXIC DIFFUSE GOITER (code No. 810-943) (Simple Goiter)

Diagnosis

There is often a history of living in an endemic area. Symptoms appear only if the enlargement is great enough to cause pressure on surrounding structures (esophagus, trachea, or recurrent laryngeal nerve). The BMR and the serum protein-bound iodine and radiiodine (I^{131}) uptake are normal.

Treatment

A Specific Measures

1. Thyroid USP B.P. 60-120 mg (1-2 gr) especially if the goiter is multinodular appears to be of value in about 50% of cases.

DIABETES INSIPIDUS (Due to Unknown Cause code No 842 770)

Destruction of the posterior pituitary or impaired function of the supraoptic nuclei or of tracts from these nuclei to the posterior pituitary (63% of cases being due to tumor) causes the condition known as diabetes insipidus. This is manifested by severe thirst and marked polyuria. A polyuria of over 6 liters per day with specific gravity below 1.006 is highly suggestive of diabetes insipidus. The diagnosis is established by the Hickey Hare test. This test consists of (1) I.V. infusion of hypertonic salt solution which in patients with diabetes insipidus causes an increase in urine flow and (2) administration of a test dose (0.2-0.3 cc) Vasopressin Injection U.S.P. B.P. (Pitressin®) which causes a decrease in urine flow.

Treatment

- A Specific Therapy Vasopressin Tannate N.N.D. (Pitressin Tannate®) 1 cc in oil I.M. is the treatment of choice. It is effective for from 24 to 72 hours. It is usually best to administer the drug in the evening so that maximal results can be obtained during sleep. Patients learn to administer the drug themselves and the dosage is adjusted as necessary. Posterior pituitary secretion inhaled 2-3 times a day may be used but it is quite irritating and absorption is uncertain. The dose varies from 30-60 mg. The aqueous preparation (Vasopressin Injection U.S.P. Pitressin®) is rarely used in chronic treatment because of its short duration of action (1-4 hours).
- B Non-specific Measures Mild cases (or Pitressin® resistant cases) require other treatment other than adequate fluid intake.
- C X-ray therapy may be used in treatment of some cases of tumor (e.g. craniopharyngeal granuloma).

THYROID

The thyroid gland utilizes inorganic iodine to form a complex physiologically active thyroxine compound that is necessary for normal body function. The normally functioning gland removes the erythrocyte concentrations of inorganic iodine present in blood synthesizes it through diiodotyrosine to thyroxine and possibly triiodothyronine and liberates the active materials probably in combination of the 2 or 3 organic compounds before excretion. There is a bound to protein. When a excess of inorganic iodine is present in the blood the thyroid cells pick it up and eliminate it from the colloid of the follicle. Under the influence of the anterior pituitary this colloid material is released with its active principle into the bloodstream. The utilization of inorganic iodine is quite constant in health ranging from 4-8 mg. per 100 cc of blood.

The requirements for iodine are very slight and difficult to estimate. About 20-200 (0.02-0.2 mg.) micrograms per day.

- 1 Patient is with severe myxedema myxedema heart disease elderly patient with hypothyroidism with other associated diseases CAUTION Begin with small doses 8-15 mg ($\frac{1}{8}$ - $\frac{1}{4}$ gr) daily for 1 week and increase dose every week by 15 mg ($\frac{1}{4}$ gr) daily up to a total of 100 to 200 mg ($1\frac{1}{2}$ - 3 gr) daily. This dosage should be continued until signs of hypothyroidism have subsided or toxic symptoms appear. Then stabilize dosage so as to maintain the B.M.R. on protein bound iodine at normal just below the level of toxicity (see below under Hyperthyroidism).
- 2 Patients with early hypothyroidism may be treated with large doses 30 mg ($\frac{1}{2}$ g) daily increasing by 30 mg ($\frac{1}{2}$ gr) every week to the limit of tolerance.
- 3 Chronic maintenance. Each patient's dose must be adjusted to obtain the optimum effect. Most patients require 50 to 130 mg (1-2 g) daily for maintenance. Optimum dosage can be determined by following protein bound iodine B.M.R. but clinical judgment is the best guide.
- 4 When required, potassium iodide sodium L-thyronine N.N.D. (Liothyronine Cytomel®) may be employed. Begin with very low dose because of its speed of action. Begin with 0.005 mg and increase slowly (see page 415).

Needless Use of Thyroid

- 1 Questionable diagnosis. If any patient can tolerate above 200 mg (3 gr) daily of thyroid the diagnosis of hypothyroidism should be questioned. No normal individual and obese individuals no hypothyroid individuals can tolerate doses up to 300 to 500 mg ($4\frac{1}{2}$ - $7\frac{1}{2}$ gr) daily without change in B.M.R. or development of toxic symptoms.
- 2 Nonspecific use of thyroid. The use of thyroid medication as nonspecific stimulant for therapy is limited only to the old method. It has been shown that the dose usually employed (100 to 200 mg or $1\frac{1}{2}$ - 3 g daily) is inefficient in altering the metabolism of normal individuals.

HYPERTHYROIDISM

- If the patient had been treated primarily to satisfy hypothyroidism according to the Graves anatomical histology of the gland as follows:
- 1 Diffuse Toxic Goiter (when associated with exophthalmos Graves disease) (see N 810 943 6)
 - 2 Nodular Toxic Goiter (see No 810 952 6)
 - 3 Hypothyroidism without Goiter (see N 810 771)
- How valuable is the treatment in determining the differential physiology and incidence of the disease that the early differential clinical and physiological thyroiditis disease and the common diagnosis of hyperthyroidism must justify.

Diagnosis

A Symptoms Nervousness irritability excitability and weight loss in spite of excessive appetite and food intake

B Signs

1 Patient must quickly all his muscles tremor and moist skin

- 2 Iodine therapy (early) If the enlargement is discovered early it may disappear completely with adequate iodine. Five drops daily of saturated solution of potassium iodide or Strong Iodine Solution U S P Aqueous Solution of Iodine B P (Lugol's solution) in $\frac{1}{2}$ glass water is adequate therapy. Continue therapy until gland returns to normal size then keep on maintenance dosage or use iodized table salt.
- 3 Iodine therapy (late) If the enlargement is of long standing, iodine therapy as above may be used but much regression in the size of the gland should not be expected.

B Indications for Surgery

- 1 Signs of pressure If signs of local pressure are present the gland should be removed surgically.
- 2 Potential malignancy Surgery should be considered for any thyroid gland with a single nodule for the chances of a single nodule being malignant are quite high. This is particularly true in younger people and when there is no response to treatment.

Prophylaxis

With an intake of 100-200 micrograms of iodine daily this condition should not occur. During times of stress (puberty, pregnancy and lactation) the upper limits of this dose may prove necessary. This amount is satisfied by 1-2 Gm (15-30 gr) of iodized salt (1:5000-1:10,000 parts iodine) daily.

HYPOTHYROIDISM (code No 810 7722)

Diagnosis

A Symptoms Early weakness, easy fatigability, cold sensitivity.

B Signs

- 1 Early These are few and may be difficult to find. Dry skin and hair, brittle nails, and menstrual disturbances are suggestive.
- 2 Later Hair tends to fall out (especially eyebrows), sweating diminishes, face becomes puffy (especially about the eyes), then non-pitting edema spreads to the rest of the body. Patient may develop anemia and heart disease.
- 3 Obesity is an uncommon finding in true hypothyroidism.

C Laboratory Diagnosis

- 1 Low BMR A BMR below -30% is suggestive but not diagnostic of hypothyroidism. A low BMR does not necessarily mean hypothyroidism; this is especially true in obese patients. (See Obesity p 380).
- 2 Serum iodine A low protein bound iodine of under 3.5-4.0 micrograms per cent (depending on the method used).
- 3 Decreased radiiodine (I^{131}) uptake (below 10% in 24 hours).
- 4 Other significant findings include elevated blood cholesterol (above patient's normal) and in severe cases anemia.

Treatment

A Specific Therapy Thyroid U S P B P is the preparation of choice. Initial dosage varies with the severity of the hypothyroidism.

drug continued the B M R will continue to fall until
the patient becomes my dermatologist

(1) The drug ppe ra to b ide 1 except f 2 fact s

(a) Drug fits reactions especially granulocytopenia. This apparently happens very infrequently with pyrimethamine and can be anticipated. The patient examined weekly or biweekly blood count taken. If the WBC fall below 4500 or if less than 45% granulocytes are present therapy should be discontinued. Other reactions are drug fever, dizziness.

(b) This is an objection of a technical nature since the gland may be mainly hyperplastic and vascular so that removal is more difficult. Because of this combined the use of a prophylactic oestrogen is probably the method of choice in preparing patients for thyroidectomy (see below).

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(b) In rare cases especially with very large glands
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A f w p t i n t m y n t p o d p c l l y t h o s w h o
h i d i o d i n e e t l y

b If th e i too lo g w it bef a gery th gi d may
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- 3 Fine tremor of extremities usually present in severe cases
- 4 Marked weight loss and emaciation
- 5 Goiter (at times a bruit may be heard over gland)
- 6 Exophthalmos may be marked
- 7 Cardiovascular findings vary most common is tachycardia but in older patients especially with long standing hyperthyroidism cardiac failure and auricular fibrillation are not uncommon

C Laboratory Findings

- 1 Elevated B.M.R. may be present in other conditions such as fever malignancy (especially leukemia)
- 2 Elevated hormonal iodine Above 8 mcg % is suggestive but this may be seen in pregnancy with excessive administration of thyroid and after therapeutic or diagnostic use of iodine containing organic compounds (e.g. drugs used for gallbladder or kidney visualization)
- 3 Increased ^{131}I uptake may be diagnostic In doubtful cases with elevated ^{131}I uptake lack of depression of uptake on thyroid or sodium iodine medication may be diagnostic
- 4 The blood cholesterol level may be low

Treatment

Treatment is aimed at stopping the excessive secretions of the thyroid. Several methods are in use and the method of choice is still open to debate and varies with each case. The most widely accepted method however is surgical preparation followed by subtotal surgical removal.

A Subtotal Thyroidectomy This is probably still the method of choice since it demands the least in follow up care. Adequate preparation is of the utmost importance. One or two drugs are generally necessary for adequate preparation iodine and/or one of the thiouracil group of drugs.

1 Thiouracil drugs (drugs of choice) Recently several thiouracil drugs or similar derivatives have been introduced. They are Propylthiouracil U.S.P. B.P. Methylthiouracil U.S.P. Methimazole U.S.P. do not contain iodine in the molecule. Iodine U.S.P. Sodium N.N.D. (Iodine ^{131}I) The modes of action of the first three are probably identical that of iodine is still not entirely clear.

a Propylthiouracil U.S.P. B.P. This drug has been most widely used and appears to be the least toxic. It is the thiouracil preparation of choice. The mode of action of this drug is such that when given in adequate dosage it prevents the thyroid gland from incorporating inorganic iodine into its organic (hormonal) form. This effect is very rapid (within a few hours) and continues as long as the drug is given. The gland often is to attempt to manufacture the hormone (resulting in hyperplastic growth becomes more so) but none is made. Because of this the B.M.R. invariably falls the rate of fall depending upon the total quantity of previously manufactured protein bound iodine available from the gland or in the circulating blood. (More protein bound iodine is present if iodine has previously been given.) The average time required for the B.M.R. to return to normal is about 4 to 6 weeks. If the

possibility of carcinogenesis (which has not yet been observed) and the possibility that an early carcinoma which might be removed surgically may remain undetected. Because of the above factors its use should generally be limited to older age group (40 or above).

D C¹³¹ Sodium Iodide Thy In the past this method was used in some select mild cases of hyperthyroidism with fair results however because of the danger of escape and because of the discovery of propylthiouracil iodine should be used only for preparation.

E X-ray Therapy Has been used in skilled hands with good results as a substitute for surgery but because of the time necessary to obtain complete effect (3-6 months) this mode of therapy should be reserved for selected cases. It is rarely indicated when 131 I is available.

F General Measures

1 Rest The patient with hyperthyroidism should be treated especially in severe cases and in preparation for surgery. With the advent of propylthiouracil mild cases are being treated as ambulatory patients. However, usually bed rest hastens recovery.

2 Diet Diet should be high in calories and protein and vitamins. These patients consume great quantities of food are generally in negative nitrogen balance and need the excess foods and vitamins because of the increased metabolic needs. Supplemental vitamin B complex should generally be employed.

3 Sedation When faced with the patients who often very nervous, sedation is always helpful and very largely qualitative may be necessary to the symptom. Phenobarbital, U.S.P. Phenobarbital, B.P. 30 mg ($\frac{1}{2}$ gr) 3-6 times daily may be necessary.

4 Testosterone propionate This drug has been shown to be of value in restoring positive nitrogen balance in these patients. May administer 25-50 mg I.M. daily or 2-3 times per week. Do not use methyltestosterone in hyperthyroidism as this aggravates the condition.

G Treatment of Complications

1 Exophthalmos The extent of exophthalmos is still unknown. Although it may be detected by the use of anteroposterior films, the evidence is still controversial. It has been shown that exophthalmos is due to edema of the extraocular muscles of the posterior chamber (muscle connective tissue). Removing the thyroid gland (by thyroidectomy or administration of propylthiouracil) does not usually help the condition and may aggravate it. It is difficult to malign exophthalmos. It has been suggested that this is due to the fact that the thyroid gland inhibits the antipituitary release of the gland allowing the antipituitary to exert more hormone and aggravate the condition. Some investigators believe that exophthalmos occurs with hyperthyroidism because the thyroid is a cation in hyperthyroidism may be equilibrated by absorption and because the basal

c It is generally impossible to bring the B M R to normal with iodine

3 Combined propylthiouracil iodine therapy The advantage of this method is that one obtains the complete inhibition of thyroid secretion with the involuting effect of iodine. This can be given in 2 ways

a Propylthiouracil followed by iodine This appears at present to be the method of choice. Begin therapy with propylthiouracil about 10-21 days before surgery is contemplated (usually B M R about +20) begin the iodine and continue for 1 week after surgery

b Concomitant administration of the 2 drugs from the start in dosages as for the individual drugs i.e. 100-200 mg propylthiouracil q i d and Strong Iodine Solution U S I Aqueous Solution of Iodine B P (L g i s s o l t o n) 10-15 drop daily

B Continuous Propylthiouracil Therapy (Medical Treatment)

Control of hyperthyroidism with propylthiouracil alone without surgery has been advocated by some

1 The advantage is that it avoids the risks and postoperative complications of surgery e.g. myxedema hypoparathyroidism

2 The disadvantage is the remoteness of the possibility of toxic reactions (see p 371) plus the necessity of watching the patient carefully for signs of hypothyroidism. Since the advent of propylthiouracil it appears that the possibility of toxic reactions is slight. The patient must report to the physician if fever, sore throat or dermatitis develop

3 Dosage

a Begin with 100-200 mg t i d to q i d and continue this dosage until the B M R is normal and all signs and symptoms of the disease have subsided then place the patient on a maintenance dose of 50-75 mg daily keeping him on the B M R or protein bound iodine to avoid hypothyroidism

b An alternative method is to continue with a dose of 50-200 mg t i d to q d. This will bring the patient to hypothyroid level keep his B M R or protein bound iodine normal with thyroid. (This may be the preferred treatment of exophthalmic goiter see page 373)

c Duration of therapy The duration of therapy and recurrence rate have not been completely worked out. However at present it would seem that of the patients kept on propylthiouracil between 8 and 18 months (the dosage slowly decreased) about 50 to 70% will show no recurrence. Increasing the duration of therapy to about 2 years or more does not increase the cure rate

C Radioactive Iodine [131] The administration of radioiodine has proved to be an excellent method for ablation of overfunctioning thyroid tissue. The rationale of treatment is that the radioiodine being concentrated in the thyroid will destroy the cells that concentrate it. Its use may be lifesaving in case of thyroid carcinoma when the cancer tissue can take up iodine. Because special techniques are not necessary to measure and handle the 131 the method is still generally limited to large medical centers. The only objections to this

hyperpyrexia tachycardia and C N S hyperirritability and delirium. The cause is uncertain but absolute or relative adrenal insufficiency may be important.

- a General treatment. Attempt to control the hyperpyrexia with cold packs and the hyperirritability with sedation.
- b Specific measures. There is no certain specific therapy. However, the use of large doses of corticotropin (ACTH) and the corticosteroids (p 424) may be life saving. The administered full doses of sodium iodide 1.2 Gm (15.30 gr) i.v. and repeated every 12-24 hours has been advocated.

DISORDERS OF CALCIUM AND PHOSPHORUS METABOLISM

NORMAL CALCIUM AND PHOSPHORUS METABOLISM

Calcium

- A Intake. Calcium is ordinarily derived from the diet. Average adult dietary intake is 0.5-0.8 Gm ($7\frac{1}{2}$ -12 g) daily and is normally adequate. During pregnancy and lactation the requirements are higher, the range being 1.5-3.0 Gm ($22\frac{1}{2}$ -45 gr) daily.
- B Absorption. Calcium is absorbed in the small intestine. Several factors influence calcium absorption:
 - 1 Vitamin D. Necessary for proper absorption.
 - 2 Presence of fatty acids or certain minerals (magnesium, potassium) may interfere with calcium absorption.
 - 3 pH of intestine. Increased acid favors absorption.
 - 4 Disease of the GI tract (e.g. chronic diarrhea, pancreatic deficiency) which disturbs motility and interferes with absorption.

Intestinal calcium metabolism

- 1 Blood. Calcium exists in plasma in 2 fractions: a diffusible fraction (45-60%) containing the ionizable active material and a non-diffusible which is bound to the globulins. When blood calcium value is reported, this is the total (diffusible and non-diffusible) calcium and may be low when protein level is low but the physiologically active ionizable portion may be normal. Therefore, always determine serum protein when serum calcium is determined.
- 2 Bone. Bone is a very actively metabolizing tissue. There is constant breakdown or resorption (osteoclastic activity) and constant new bone matrix formation (osteoblastic activity) and constant calcifying of this matrix. The activity of building pathologic bone (osteoblastic activity) is associated with the presence of alkaline phosphatase enzyme and its liberation into the blood. This enzyme is increased when osteoblastic activity is increased.
- 3 Excretion. Calcium is excreted in the urine and stools. Most of the stool calcium is derived from unabsorbed dietary calcium and varies with calcium intake and absorption. The urinary calcium varies with the amount absorbed but the variation is not as great as with the stool calcium.

secretion does not have any pituitary depressing effect. Therefore it would seem rational to treat this condition by giving thyroid orally.

- a Thyroid dosage. Immediately after surgery or after B M R has returned to almost normal ($+20\%$) with propylthiouracil therapy begin giving thyroid 100-200 mg daily. Give dosage adequate to maintain B M R at about $+20\%$. This therapy should be used whenever there is a tendency for progression of the exophthalmos although it is not always effective.
 - b Physical protection of eyes. Dark glasses protect from dust eye shields tarsorrhaphy and other measures may be necessary. Ophthalmological consultation should be requested.
 - c Corticotropin (ACTH) or corticosteroids. The use of these agents in large doses has been proposed. In some cases they have proved helpful. They probably act by reducing the inflammatory reaction which occurs in the periorbital tissues.
 - d Surgery of malignant exophthalmos. Every patient with exophthalmos should have actual and periodic measurements made with an exophthalmometer. One should not rely upon clinical judgment to determine whether or not exophthalmos is present or progressing. In severe progressive cases where corneal edema limitation of extraocular muscle movements and failing vision occur it becomes practically a surgical emergency to save the eyesight. The operation of choice is orbital decompression.
- 2 Cardiac complications. Whether or not thyrotoxicosis is itself can cause heart disease is still unsettled however a number of cardiac complications are at times associated with hyperthyroidism.
- a Tachycardia. Some degree of tachycardia is always found if normal rhythm is present in thyrotoxicosis. This requires only the treatment of the thyrotoxicosis.
 - b Congestive failure. This tends to occur in longstanding thyrotoxicosis especially in the older age groups. Therapy is the same as for congestive failure from any cause. Digitalis seems to be effective in congestive failure associated with thyrotoxicosis (see p 197).
 - c Atrial fibrillation. May occur in association with thyrotoxicosis. Treat as a symptomatic arrhythmia but do not try to convert the atrial fibrillation in a toxic patient. Most cases will revert to normal rhythm soon after toxic symptoms subside. However if fibrillation remains for 2 weeks after surgery or for 2-4 weeks after B M R has returned to normal using propylthiouracil therapy one should consider use of quinidine to convert to a normal rhythm (if no contraindications are present) (see page 200).
- 3 Thyroid storm. Fortunately this condition is rare with modern forms of therapy. It occurs now mainly with inadequately treated iodine deficiency states immediately after subtotal thyroidectomy. It is characterized by

laid down in adequate amounts however the matrix that is deposited is calcified normally

Blood

A Calcium Disorders (Normal values 9-11 mg %)

- 1 Hypocalcemia Observed in a number of abnormal conditions. If severe it may result in:
 - a Hypoparathyroidism
 - b Osteomalacia if severe of whatever etiology (e.g. that associated with steatohepatitis resulting in improper calcium absorption)
 - c Due to phosphorus retention by the kidney (e.g. nephritis) which causes an elevated serum phosphorus
- 2 Hypercalcemia
 - a Hyperparathyroidism
 - b Multiple myeloma or other diseases with high protein
 - c Overdose with dihydrocholesterol (A.T. 10) or vitamin D

B Phosphorus Disorders (Normal adult values 2-4 mg %)

- 1 Hypophosphatemia Occurs in hyperparathyroidism and may occur in osteomalacia but low serum phosphorus values are rare. A rare cause of hypophosphatemia is the de Toni-Debré syndrome
- 2 Hyperphosphatemia Serum phosphorus level is elevated in growing children and in a hemodialyzed patient. The high serum phosphorus levels are found in renal failure which causes phosphorus retention

HYPERPARATHYROIDISM

(Adenoma, code No 820 8046A) (Hyperplasia, code No 80 943 6)
(Osteitis Fibrosa Cystica Generalized, code No 200 773)

An increase in parathyroid hormone leading in phosphorus and calcium from the body. An x-ray and polyuria are evidence of bone loss. This is found on x-ray and shown by the presence of pathological changes. Clinically all of the following are found: hypoparathyroidism, low serum phosphorus, high serum calcium, low urinary phosphorus, high serum phosphorus, and increased tubular reabsorption.

Treatment

A Surgery If parathyroid tumor is usually found. It should be removed surgically.

CAUTION After surgery patient may in the course of several hours develop tetany as a result of rapid fall of blood calcium. The level may only fall to the normal range but due to the rapid alteration in level tetany may be precipitated. The possibility of hypoparathyroidism (see below)

B Fluids Large fluid intake is necessary so a diluted urine will be excreted and minimize the formation of renal calculi from serum phosphate

Phosphorus

Phosphorus is involved in many metabolic reactions and in some of these it acts with calcium. In general there is an inverse ratio between calcium and phosphorus in the blood. When calcium is high phosphorus usually is low and vice versa.

- A Intake Phosphorus is a rich constituent of many food stuffs and shortages are unknown except in starvation.
- B Absorption Most phosphorus is absorbed as simple inorganic phosphorus.
- C Intermediary Metabolism (Blood phosphorus level is usually from 2 to 4 mg % in adults)
 - 1 Bone Phosphorus moves with calcium into bone to form calcium phosphate salts.
 - 2 Phosphate ion (PO_4) is very important in many metabolic reactions and the potential energies of these phosphate bonds especially with carbohydrate are the sources of much of the entire body energy. In carbohydrate metabolism PO_4 forms hexose phosphate bonds and in fat metabolism it acts through the phospholipids. It also participates in nucleic acid metabolism.

ABNORMAL CALCIUM AND PHOSPHORUS METABOLISM

Bone

Most abnormal calcium and phosphorus metabolism is concerned with abnormal osteogenesis. Therefore to understand the therapy of the disorders of calcium and phosphorus metabolism one must understand the abnormal physiology and the physiological differences between the metabolic diseases of poorly calcified bones.

- A Osteomalacia A disorder of calcium or phosphorus metabolism of such nature that there is absence of or poor deposition of calcium salts in the normally formed osteoid matrix. The weakened bone therefore is subjected to many mechanical stresses which act as stimulants to the osteoblasts to lay down more osteoid matrix. This increased osteoblastic activity leads to an elevated serum alkaline phosphatase. The serum calcium and/or phosphorus may be depressed or normal. The product of calcium times phosphorus is generally decreased.
- B Osteitis Fibrosa Cystica or Cystica A disorder of calcium or phosphorus metabolism in which there is increased resorption of bone due to hyperparathyroidism. The excessive parathyroid hormone probably has a dual action: it diminishes renal tubular reabsorption of phosphorus causing excessive phosphorus loss and calcium phosphate is mobilized from bone raising serum calcium which then spills over in urine. There is probably direct stimulation of osteoclasts by the hormone. Since the bone is weakened the remedy may be added to ease strains as in osteomalacia which may stimulate osteoblastic activity and cause a high serum alkaline phosphatase. This results in a high serum calcium, a low serum phosphorus and a high calcium and phosphorus excretion in the urine.
- C Osteoporosis Strictly speaking osteoporosis is not a calcium or phosphorus metabolic disorder. It is due to a decrease in the osteoblastic activity and hence the protein matrix is not

phosphorus and help restore normal calcium level to normal in this condition.

- 6 Aluminum hydroxide gel may be employed to help with the serum phosphorus (see p 302)

OSTEOPOROSIS (Senile Osteoporosis code No 200 798)

Osteoporosis occurs most commonly in postmenopausal patients. It is also found associated with other conditions leading to generalised osteoporosis e.g. malnutrition particularly due to low protein intake. Cushing's syndrome and excessive use of corticosteroids (ACTH) or other corticosteroids atrophy (where stimulation of osteogenesis is absent) commonly may be involved.

The first complaint is usually backache. There is x-ray evidence of rarefaction especially in the lumbar vertebrae and pelvis and often a collapse and fracture of vertebral bodies. Other pathological fractures may occur. The blood alkaline phosphatase and alkaline phosphatase are normal.

Treatment

A Specific Measures Vary with the cause but combined hormone therapy usually indicated.

- 1 Postmenopausal (mostly in females)

Estrogen may be effective in stimulating osteoblasts before beginning estrogen therapy in a postmenopausal woman perform careful pelvic examination to rule out neoplasia or other abnormality and warn patient of a relatively high vaginal bleeding may occur. Administer estrogen daily except first 5 or 7 calendar days of a 4 month and then repeat cycle.

(1) Diethylstilbestrol U.S.P. Stilboestrol B.P. 0.5 2.0 mg daily as tolerated.

(2) Ethinyl Estradiol U.S.P. 0.02 0.05 mg daily as tolerated.

- b Testosterone For its protein anabolic effect and hence its tendency to lay down bone matrix testosterone may be used in addition to estrogens. Give 10-20 mg methyltestosterone orally. Avoid overdosage in female excise embryonic appearance of male secondary sexual characteristics. However, these will regress if therapy stopped.

- 2 Old age and idiopathic As for postmenopausal both testosterone and estrogen should be used in both males and females.

- 3 Protein with malnutrition Adequate diet is of great importance. However, the hormone may be used as before if response to diet is poor.

- 4 Cushing's disease (see p 383)

B General Measures

- 1 Diet Should be high protein high calcium (milk and milk products desirable). Vitamins supplement especially vitamin D 2000-5000 unit daily may be given also.
- 2 Activity Patient should be kept as active as is indicated by use of tiludronate.

HYPOPARATHYROIDISM (Unknown Cause code No 820 x10) (Injury due to Operation code No 820-415.x)

A deficiency of parathyroid hormone usually occurring post operatively following thyroidectomy or surgery for parathyroid tumor or hyperplasia. It is characterized by muscle weakness, irritability, tetany, a low blood calcium, high or normal blood phosphorus, normal phosphatase, and normal bones by x ray (except after removal of parathyroid tumor). Cataracts may occur particularly in young persons.

Treatment

- A Emergency Treatment for Acute Attack (Hypoparathyroid Tetany) Usually postoperative and requires immediate treatment
1. Calcium Chloride U S P B P 5-10 cc (1 2 1/2 dr) of 10% solution I V slowly until tetany ceases or Calcium Gluconate Injection U S P B P 10-30 cc (1/3 1 oz) of 10% solution I V may be given. 10-50 cc of either solution may be added to 1000 cc of 5% glucose in water or saline and administered by slow I V drip. The rate should be so adjusted that hourly determination of urinary calcium by means of the Sulkowitch test will be positive.
 2. Parathyroid Injection U S P 50-100 units (1/2 1 cc) I M or s.c. 3-5 times daily as necessary to prevent tetany. Do not use parathyroid hormone for over one week because refractory state tends to develop rapidly. Use only as long as absolutely necessary. A usually parathormone is rarely ever used. It is not very practical and usually not necessary.
 3. Calcium salts should be given orally as soon as possible. calcium gluconate 4 Gm (60 gr) q i d calcium lactate calcium chloride 2-3 Gm (30-45 gr) q i d
 4. Dihydroxycholesterol N N D (Hytakerol®) should be given as soon as oral calcium is begun. Begin with 4-10 cc (1 2 1/2 dr) of oily sol (1-25 mg per cc) orally daily for 2-4 days then reduce dose to 1-2 cc daily for 1-3 weeks and then determine maintenance requirements.
- B Maintenance Treatment
1. High calcium low phosphorus diet (milk milk)
 2. Calcium salts as above may be continued (except chloride)
 3. Dihydroxycholesterol N N D (Hytakerol®) 1/2 1 cc daily or 3 times weekly to maintain blood calcium at normal level
 4. Calciferol U S P 1-5 mg daily. In some cases up to 7 or 8 mg calciferol daily may be substituted for dihydroxycholesterol. Vitamin D action is probably similar to that of dihydroxycholesterol and it can certainly be substituted adequately clinically. The initial action of vitamin D appears to be slower. However, the cost to the patient is less than using dihydroxycholesterol and the margin of safety is probably greater. Regulate the dose by daily Sulkowitch test which should run 1-2+.
 5. In some patients Probenecid N N D (Benmid®) in doses of 2-4 Gm (30-60 gr) daily has been shown to block the tubular reabsorption of phosphate and hence to lower serum

TreatmentA Specific Therapy (Chronic Cases)

- 1 Cortisone or hydrocortisone. The drugs of choice at present. Most Addisonian patients are well maintained on 6-25 mg ($1/10$ - $3/8$ gr.) daily given in divided doses 3 to 4 times daily orally. On this dose most of the metabolic abnormalities are corrected. Most patients, however, do not obtain sufficient salt-retaining effect and require D O C A or fludrocortisone supplementation or traditional salt.
- 2 Desoxycortosterone Acetate U S P. D oxycorton Acetate B P (D O C A). This drug controls electrolyte balance and has no other significant metabolic effect. Intramuscular administration of D O C A may be used initially but is rarely necessary. The usual dose for replacement is 1-4 mg daily. When the response has been adequate (see below) change to buccal use.
 - a Buccal use of D O C A. One tablet daily or at most 1 tablet twice daily will give adequate replacement. The tablet is placed between the cheek and teeth and allowed to dissolve (see p. 418). Desoxycortisone dimethyl acetate 25-75 mg I M once monthly may be substituted for D O C A (25 mg I M once monthly but 1 mg D O C A in 11 per day).
- 3 Fludrocortisone Acetate N N D. This new drug is very potent in inducing sodium retention. It is effective orally. Dose is 0.1 to 0.25 mg daily or every other day (see p. 424).

CAUTION: Whenver using D O C A or fludrocortisone, avoid excessive sodium. Do not have patients on low potassium diets when giving these drugs for the first time; development of potassium deficiency.
- 4 Sodium chloride in large dose (5-20 Gm. daily) may be used to supplement corticosteroid in the balance of D O C A or fludrocortisone.

B General Measures

- 1 Diet. Give high carbohydrate high protein diets. Frequent small feedings tend to be better tolerated than 3 large ones.
- 2 Avoid exposure to infection and to tall infections immediately and vigorously.
- 3 Methyldopa 10-20 mg daily orally or 10-20 mg I M 3 times weekly is often helpful for its protein-anabolic effect and for the hypotensive effect of well-being it induces in the debilitated patient.

Criteria of Adequate Therapy and OverdosageA Adequate Therapy

- 1 Return of blood pressure to normal. May require 2 to 3 months with adequate therapy.
- 2 Maintenance of normal fasting blood glucose level.
- 3 Return of plasma electrolyte to normal level.
- 4 Weight gain (usually due to fluid).
- 5 Improvement of appetite and of general strength.
- 6 Increase in level of heart to normal.

B Overdosage. Must be watched for and avoided very carefully especially in patients with cardiac or renal compensation.

- 1 Signs and symptoms of corticosteroid overdosage (see p. 423).
- 2 Development of deepened moon face or excessive weight gain.

OSTEOMALACIA (code No 200 7642)

Osteomalacia results from calcium deficiency due to any cause. In adults these include vitamin D lack or resistance (rare) and sprue syndrome or pancreatic disease. Chronic renal disease may also produce osteomalacia but more generally produces secondary hyperparathyroidism.

Rickets (code No 010 764) is the childhood type due to inadequate intake of vitamin D. Laboratory findings include a low or normal serum calcium or a low or normal serum phosphorus (except in renal disease where it may be elevated) and most characteristically an elevated phosphatase.

Treatment

A Specific Therapy

- 1 Rickets Vitamin D even in small doses is specific. 2000-5000 units daily are adequate.
- 2 Adult osteomalacia and Milkman's syndrome Vitamin D is specific but very large doses are necessary to overcome the absorption defect. Give until an effect is noted on blood calcium. Usual dose is 25 000-100 000 units daily. Doses up to 300 000 units daily may be necessary but if the doses are over 100 000 daily they must be used cautiously.
- 3 Pancreatic insufficiency (see p 289). Adequate replacement therapy is of paramount importance. High calcium intake and vitamin D 2000-10 000 units daily are also of value.
- 4 Sprue syndrome Folic acid and vitamin B₁₂ appear to be of value (see p 226).
- 5 Some are forms of renal disease. Treatment is aimed at the altered renal physiology.

B General Measures. High calcium diet and calcium gluconate, calcium lactate or calcium chloride 5-20 Gm (1-5 dr) daily.

ADRENAL CORTEX

ADDISON'S DISEASE (Adrenocortical Insufficiency)

(Due to Tuberculosis code No 880 123.x)

(Undetermined Cause code No 861 782)

A disease due to lack of secretion of adrenal cortex caused by tuberculous destruction of the gland, surgery or undetermined factors. It is manifested by asthenia, anorexia and other GI disturbances, hypotension and pigmentation usually brownish of the skin and mucous membranes. This pigmentation is mainly an accentuation of already pigmented areas and a deposition of pigment in skin creases.

The laboratory findings include a low blood sugar, increased insulin sensitivity, low blood sodium and chloride, elevated potassium, elevated N/P/N and a positive water test. There is a decrease of 17-ketosteroid and corticosteroid excretion and lack of response to adrenocortical tropic hormone in primary Addison's disease.

1. Ovary: Ovulation usually delayed to odum re-
te tio my es l t in e ebr l dema (with un scousness
o o vuls ons) o p lmona yed m Withhold od m a d
fl id t mpo arily and tr at fo th conditi ns (s e p 15)
2. Hypokalemia: Flaccid paralysis with lws un K^+
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t eated w th K^+ salts (s p 15)
3. Hypopyla: This complicat on ra e w th p ese t tr t
me t m th ds (S e p 27 for ther py)
4. For oth r ompli t os of dr al st rod th apy (e g
psych ti re t os) s p 4 8

CUSHING'S SYNDROME (Adrenocortical Hyperfunction)
(Pituitary Basophilism code No 841 7763)
(Adrenocortical Hyperfunction code No 861 7813)

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- 3 Development of hypertension
- 4 Increase of diameter of heart above normal
- 5 Development of signs of potassium deficiency (weakness followed by loss of muscle power and finally paralysis) especially if patient is on a low potassium diet

Treatment of Adrenal Crisis (Acute Adrenal Insufficiency)

The adrenal crisis is an emergency. The patient must be treated vigorously and observed constantly until well out of danger. **Overtreat rather than undertreat.**

A Severe Crisis

1 Emergency treatment

- a Anti shock measures. Use appropriate adjunct measures (see p. 27) especially plasma vasopressor drugs and oxygen. Do not use narcotics or sedatives.
- b Hydrocortisone. Administer hydrocortisone hemisuccinate (Solu Cortef®) 100 mg dissolved in 2-10 cc water I.V. over about 1 minute. Follow with hydrocortisone free alcohol (Infusion Concentrate Hydrocortisone® Cortef® Sterile solution) or Solu Cortef® 100 mg in 1 L 5% glucose in physiological saline solution by I.V. infusion over a period of 2 to 8 hours. If Solu Cortef® is not available initially administer the hydrocortisone free alcohol 100 mg in 1 L of 5% glucose in physiological saline I.V. over a period of 1 to 3 hours. An additional 50-100 mg of hydrocortisone may be added to subsequent infusions during the first 24 hours if necessary. (If parenteral hydrocortisone is not available give aqueous adrenal cortical extract 25-50 cc I.V. immediately and follow with 100-200 cc aqueous adrenal cortical extract in 1000 cc saline dextrose infusion.)
- c Cortisone acetate. Give initial cortisone acetate 10-20 mg I.M. in four different sites (total 40-100 mg). Follow with single injections of cortisone 25-50 mg I.M. every 6 hours and gradually lengthen intervals of administration to 25 mg every 8 hours.
- d Subsequent parenteral fluid. After the first I.V. infusion mentioned above is completed it should be followed immediately by 1000 cc 10% glucose in physiological saline solution (including additional hydrocortisone as mentioned above). The total fluids in the first 24 hours and daily thereafter should not be greater than 3 liters in order to avoid excessive administration of sodium.
- e Antiseptic measures. If infection is present treat intensively with indicated antibiotics.

2 Convalescent treatment When patient is able to take food by mouth place on oral cortisone 12.5-25 mg every 6 hours and reduce dosage to maintain the levels as needed.

B Moderate Crisis If the patient's physical condition does not appear to be critical and is not associated with a significant degree of shock the treatment outlined above may be modified by appropriate reduction in dosage. However, it is generally best to overfire the patient in moderate crisis during the first 24 hours rather than ask undue treatment.

C Complications Arising During Course of Treatment of Crisis

MALE HYPOGONADISM

Etiology

Failure of the gland may result from several causes. This failure may be primary (due to testicular disease) or secondary to malfunction of other glands, most often the pituitary or thyroid.

Diagnosis

The physiologic diagnosis of the etiology of hypogonadism (e.g., primary or secondary) is usually based on laboratory tests.

Type of Hypogonadism	Urinary 17 K test results	Urinary Gonadotropins
Primary	Low or normal	Elevated
Secondary Pituitary	Usually low but may be normal	Very low
Androgenic Neurasthenia	Low or normal	Low or normal. Not generally as low as pituitary type
Thyroid (Thyroid?)	Low or normal	Low

Many clinical syndromes have been described but basically they all fall into one of two categories. The difference is noted below.

A. Primary Type Should not be diagnosed before 18 years of age. This is a failure of development of normal testicular function and is manifested by small or absent testes, small penis, lack or diminution of secondary and pubic hair, lack of masculine development of the young boy, and the chronologic age. Some of the patients (e.g., those with Klinefelter syndrome) have been known to have female characteristics.

B. Secondary Type Loss of function of normal testicular function (e.g., after trauma and infection, and malnutrition). Apparently the normal testis of testicular function, the male is not generally as sensitive and to maintain in life the function of ovarian function. The female hormone is often missing, may be a therapeutic effect with the development of symptoms as in female menopause. (p. 388)

Treatment

Testosterone (the male sex hormone) is the drug used for replacement therapy in the male. (For a parathyroid available see p. 420.)

A. Primary Hypogonadism. Adequate treatment with therapy can

continue the I M and possibly the oral dosage during and after surgery. After surgery gradually decrease the dose and maintain the patient as a chronic addisonian patient (see p. 381). Because of danger of precipitating heart failure care must be exercised to avoid excessive fluids and sodium.

2. X-ray therapy to the pituitary may be of value only in rare cases of hyperplasia.

B. General Measures

1. Diet: High protein diet should be given although dietary attempts to correct the negative nitrogen balance are doomed to failure.
2. Hormones:
 - a. Testosterone has been of value in reversing the negative nitrogen balance. For this testosterone propionate in oil 25-50 mg daily I M has been found necessary.
 - b. Inulin is usually of little or no value in controlling the glycosuria and hyperglycemia. Insulin is usually unnecessary as the diabetes is quite mild.

VIRILIZING DISEASES OF FEMALES

(Due to Tumor code No. 8041)

In the adult the virilizing or masculinizing disease is usually caused by a tumor arising in the adrenal ovary or from cell rests of one of the above tissues. It is characterized by excessive hirsutism (especially of male type), amenorrhea, enlargement of clitoris, deepening of voice, excessive musculature, and excessive 17 ketosteroid excretion. Surgical removal of the tumor is the treatment of choice.

Another form of the disease begins in childhood. It is due to overproduction of androgen type hormones from bilaterally hyperfunctioning adrenal cortices. In many of these patients there may be associated manifestations of hypoadrenocorticism (i.e., excessive salt and water loss and failure to maintain a fasting blood sugar). Treatment with corticoids has proved valuable in reducing the activity of the glands (particularly through suppression of endogenous ACTH) and in supplying exogenously needed corticoids. In adults the drugs of choice appear to be prednisone or prednisolone in doses of 5-25 mg daily orally in divided doses. Some investigators feel that the same dose of cortisone acetate by the I M route may be more efficacious in this syndrome.

Most cases of excessive hirsutism in females are not due to endocrine disease but to hereditary or racial factors and should not and cannot be treated by any internal medications or surgery.

HYPOGONADISM

Hypogonadism is due to a failure of the sex glands to elaborate sufficient quantities of their hormones to bring about or maintain the appearance of secondary sexual characteristics.

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make these individuals into normal adult males in every way except that they cannot produce sperm. These patients must be placed on testosterone and maintained for life on adequate doses of testosterone. There is little evidence that any pituitary substance or gonadotropin is of significant value in treating these patients (see pp 412-413).

- 1 Long acting testosterone preparations 200-500 mg IM every 2-4 weeks may be employed (see page 420)
- 2 Free testosterone pellet implantation in experienced hand is an excellent method. The dosage is as follows 300 mg is the minimal but an effective dose. Average is about 600 mg. For maximum effect about 900 mg are implanted. The pellets are implanted subcutaneously with a trocar or into a pocket made by blunt dissection. Either the infra-scapular area or anterior thigh area is used. Testosterone pellets remain and are effective for 3-4 months and then must be replaced.
- 3 An alternative method is the oral or sublingual administration of the drug to the patient and entails all the difficulties of prolonged oral administration. Dosage varies with various individuals but 10-25 mg daily orally is usually adequate dosage to cause maturation and virilization and maintenance of this state. Evidence now dictates that there is no advantage of buccal over oral administration.

B Postpubertal Oral use of methyltestosterone is probably the method of choice. The dosage necessary to control symptoms and to aid in overcoming the protein loss and debility of age is often as low as 5-20 mg daily. This dose may be used for a short period of time to control symptoms or may be continued indefinitely for control of symptoms and for its protein anabolic effect. The long acting testosterone by injection may also be employed.

FEMALE HYPOGONADISM

The most common symptom of female hypogonadism is amenorrhea. However most cases of late amenorrhea are not due to hypogonadism.

AMENORRHEA (code No. 761)

Etiology

Normal menstruation depends on many factors. From the functional point of view there must be an intact pituitary gonadotrophic axis. Any break in the cycle of production of the various pituitary or ovarian hormones concerned with normal menstruation or the lack of an endometrium capable of response to ovarian hormones will result in amenorrhea. Treatment is based on determining the measure on the level at which the disturbed physiology exists.

- a If pati nt has regular period she probably is manufac turing suff c t estrog and does not n ed th rapy
- b If cycl are very irregula and the patient suff rs f om m nopausal symptoms t ogens given in cy li al fa hion may be helpful Begin estrogens about 5 d ys afte onset of last menstrual period and ontinue in a cy lic fa hion Ethinyl Estradiol U S P 0 05 mg o Di thylstilbestrol U S P Stilboe t 1 B P 0 5 mg by mouth daily exc pt for the fi st 5 day of e ch month This is imple for patients to rem mber
- c If patient has b com am no h l there is no reas n to give troge in do la ge enough to reinstitute menses b t only to cont l symptom
- d Duration f therapy Th ha not been standard d and m st b adjust d to the individual ca Thr months to 1 y r usually suff ces
Maint na dos for lif Be au of the anaboli ffect of estrog a d b use of th r known beneficial eff t on cal ium m tabolism trogen th rapy ha b n recom m nded f r the d rati n of lif fo women beyond th menop use The advisab lity of this p ti e r mains un s tled

If a pati nt i on long term t og n therapy sh should ke pan ac t record of h dosag bedul a d bl ding Wh never bleeding occ rs that is not on sch dule (d ring w thdrawal pha) an inv tigation f tumor should be in tit ted V ginal cytologi al examina tion should b rri d o t routin ly v ry 6 m ths or 1 y

- 2 P y hologi l spe t M y of the ymptom of th meno paus are undo bt dly p y hologi al Th most ommon symptom i nxi ty more s v emotional disorde s may occu

a S dation Ph obarb tal U S P Ph nobarb tone B P 15 mg (1/4 gr) t i d q i d

b P y ho th rapy Simpl xplanation and re s ran e that th liv n d n t b hanged be a s of th menopause a ad quat in m st patient In th m e ev re cas s howev th x d of p y hiatrist m y prove sary

B S gi l nd X ray M opa Th s s diffe from the nat ral m pa only in th b uptness and verity of the sympt m both phy i l gi al and p ych l gi al In these p ti nt it i advisab t help th pati nt liv as n rmal a life as po bl If th pati nt an be mad to hav normal periods and if th y und tand that thei m trual y l will go on un ha ged th y usually mak tabl adjustments Estrogen th rapy i a f natural menopause (above)

Compli tions of Postm nopaus l Hypogonadi m nd Tr tment

A Ost poro (P 378)

B Se il V ginal G v Di thyl tilbe t ol U S P Stilboe trol B P 0 1 0 5 mg o oth tr gen (s p g 422) daily

ally Stilbe t ol vaginal upposito i co taini g 1 mg may be used daily f 10 t 14 days while continui g oral tilbestrol

not (see p 413) Employ estrogen alone or in combination with progesteron (see p 422)

(2) With high urinary gonadotropins Gonadotropins are likewise of no value treat as above

2 General measures

- a Adequate diet and dietary treatment as needed to correct abnormal deviation of weight
- b Psychotherapy in cases of emotional disturbances
- c Correction of anemia (see p 219)
- d Correction of any other metabolic abnormality (e.g. hypothyroidism)

MENOPAUSE

Etiology

Failure of the ovaries with cessation of menstruation may result from several causes most commonly the natural menopause and menopause due to surgical removal or x ray Failure may be secondary to malfunction of other glands usually pituitary or thyroid

A Menopausal Syndrome (code No 805)

B Artificial Menopause Due to Roentgen Rays (code No 788 471)

C Hypofunction of Ovary Due to Unknown Cause (code No 788 x10)

Diagnosis

The menopause is due to a loss of ovarian function and is manifested by vasomotor disturbances (e.g. hot flashes) nervousness emotional instability and amenorrhea Abnormal uterine bleeding may occur in the natural menopause before amenorrhea sets in

Treatment

A Natural Menopause The menopause in the female is characterized by at least 2 important factors *physiological* failure of ovarian function which occurs rather rapidly in the female and *psychological* recognition of the fact that reproductive life is at an end Many believe that a measure of this there is a marked change of life an implication that in addition to cessation of reproductive function there is complete alteration in one's way of life sexual activities personal relationships etc This latter belief is entirely erroneous Most women go through the menopause with out any difficulty in fact without symptoms However in those having symptoms one must carefully evaluate the role of the physiological and psychological factors before beginning any therapy Most cases will have a mixture of physiological and psychological factors

- 1 Physiological aspects (estrogen therapy) The characteristic symptoms that seem undoubtedly to be due to the cessation of ovarian function the most prominent being vasomotor instability (e.g. flushing) Another may be the feeling of tension especially in the head In women who suffer primarily from symptoms of cessation of ovarian function use of estrogens is indicated The dosage and method of administration used depends on whether or not the patient is still menstruating

MENORRHAGIA (code No 785 x20) FUNCTIONAL UTERINE BLEEDING

With the gradual failure of ovarian function excessive bleeding at the time of the menses is common. This is due to prolonged hypoestrogenic effect without concomitant progesterone production.

Another type of functional uterine bleeding which occurs most commonly in young women is due to a hyperestrogenic effect. In any case of prolonged and unusual bleeding suspect and rule out neoplasms of the uterus.

Treatment consists of administration of progesterone 100 mg orally or 10 to 15 mg I.M. daily for 5 days or 125 to 250 mg Hydroxyprogesterone Caproate N.N.D. (Delalutin®) I.M. once. This converts the proliferative endometrium into a secretory one with complete shedding when the progesterone is withdrawn.

OBESITY (code No 007)

(Due to Excess Food code No 010 70x)

(Undetermined Cause code No 010 70y)

Obesity may be defined as an increase in weight of over 10% above normal due to generalized deposition of fat in the body.

Normal weight is difficult to determine for average clinical practice. However, normal as defined by the standard age, height and weight tables is satisfactory (see standard body chart).

From a metabolic point of view, all obesity has a common cause: intake of more calories than are required for energy metabolism. The reasons for differences in the energy utilization of various individuals whereby that one person can utilize his calories more efficiently than another is unknown. Although most cases of obesity are due to simple overeating, a few endocrine disorders lead to specific types of obesity (e.g., Cushing's syndrome and hypothalamic lesions) but these conditions are rare. Hypothyroidism on the other hand is not necessarily associated with obesity.

Treatment

Specific weight reducing chemical agents and hormones singly or in combination are either ineffective or hazardous and have no place in the treatment of exogenous obesity.

A Diet (see p. 58) Diet is the most important factor in the management of obesity. There are a number of points to consider:

1. Calories In order to lose weight it is necessary to decrease the intake to below the caloric requirement of the individual (see p. 47). One can determine a very approximate average weight loss per day with a given diet by the following formula:

$$\frac{\text{Approximate Caloric Requirement Per Day} - \text{Number of Calories in Diet}}{4000} = \text{Weight Loss in lbs./Day}$$

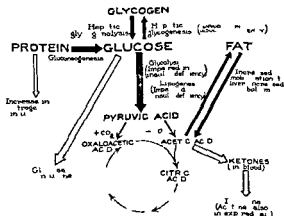
The number of calories per day to prescribe for a patient varies with age or up to a certain temperament and the urgency of the need for weight reduction. A daily caloric intake of 800-1200 Calories is satisfactory. ing diet

the administration of insulin. Although failure of the islet cells of the pancreas to manufacture sufficient insulin to supply the endogenous demands of the body is an important factor, this is not the only factor in clinical diabetes. The best evidence for this is the small insulin requirement of totally depancreatized humans as compared with the insulin requirements of patients with spontaneous occurring diabetes.

Physiological Physiology

The primary metabolic defect in diabetes appears to be an inability to metabolize glucose properly. However, this defect has a marked secondary influence on protein and fat metabolism. The complicated disturbed metabolism can best be shown by oversimplified diagrams. The sketch on p. 392 shows the normal interrelationships. The presumed site of action of insulin is indicated.

However, in diabetes, since insulin is diminished or absent, other metabolic pathways, all of them catabolic, take precedence. The following diagram shows the relationships in diabetes.



Abnormal Glucose Metabolism in Uncomplicated Diabetes (Report of the Committee on the Nomenclature of the American Diabetes Association, 1957, Lang Medical Publication, Los Angeles, California)

From the above diagram it can be seen that as the result of impaired glucose oxidation the following major metabolic alterations have occurred:
1. Glucose is lost into the urine, diminishing the body's store of carbohydrate.

weight for fat tissues have a definite but slow metabolism. It has been shown that obese people with low B M R's can tolerate 0.2 Gm (3 gr) or more of thyroid per day without change in B M R. Prolonged administration of thyroid may suppress the patient's normal thyroid secretion.

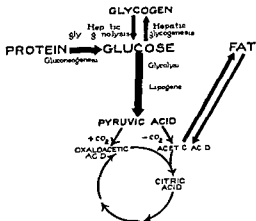
b *Dinitrophenol* must never be given to any patient.

It is a toxic drug that may cause blindness or even death.

- C Exercise Although exercise increases the energy output, extreme exercise is necessary to significantly alter weight. Playing 18 holes of golf for instance raises the total caloric requirements only by about 100-150 Calories. In addition, exercise tends to increase the appetite and may make it more difficult to control the diet properly.
- D Psychological Factors Overeating is largely a matter of habit and in some cases may be associated with deep underlying neurosis. Whatever the cause, the patients must be retrained in their eating habits and educated to understand that once their weight is normal, they can easily become obese again by eating more than their normal caloric needs.

DIABETES MELLITUS (code No. 871 785)

Diabetes mellitus in man is a metabolic disorder of unknown etiology, the metabolic defect of which appears to be corrected by



Normal Interrelationship in Glucose Metabolism

I H

Insulin is utilized clinically to enhance carbohydrate oxidation. This is measured clinically by noting the lowering of the blood sugar or the disappearance of glycosuria.

A Duration of Action of Insulin Preparation. There are 3 main types of insulin available:

1. Short acting insulin: Insulin Injection U.S.P. B.P.
Regulin insulin

b. Crystalline zinc insulin

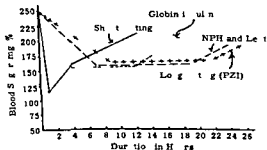
(For clinical purposes the actions of these 2 insulins are identical. Crystalline insulin is prepared only because of its greater purity). These are useful mainly in controlling postprandial blood sugar elevations.

2. Long acting insulin: Protamine Zinc Insulin Injection U.S.P. B.P. This is useful in lowering the mild or hypoglycemia which is present during the main part of the time between meals.

3. Intermediate acting insulin

Isophan Insulin Injection U.S.P. (NPH). A stable mixture with protamine which is a 21 minute delayed action insulin. It may also be tailored to fit the patient by addition of appropriate amounts of regular insulin.

b. Lente Insulin N.N.D. Made by a ratio of zinc to insulin and proportions (protamine and phosphate free). Action is almost identical with that of NPH insulin. Globulin with Zinc U.S.P. This insulin is similar in action to 21 minute insulin but that is delayed of effect as compared. It is used in many diabetic patients but it cannot be mixed with short acting insulin. Intermediate insulin. See page 395.



Extent and Duration of Action of Various Insulins
(in a Fasting Diabetic)

- 2 Liver glycogen stores become depleted in supplying glucose to the body
- 3 Protein stores begin to break down at an excessive rate to supply glucose (gluconeogenesis)
- 4 Fat mobilization to the liver and catabolism are increased
- 5 The excess acetate formed cannot be immediately utilized and condenses to give rise to ketone bodies (β -hydroxy butyric acid and acetoacetic acid)
- 6 As a result of increased ketone body formation and the slow utilization ketone bodies accumulate in the blood and spill over into the urine. Since these substances are acids they are excreted in the urine joined to fixed base (Na^+ , K^+ , Ca^{++} , etc.). The accumulation of acid ions and loss of fixed base results in the condition known as acidosis.

Diagnosis

The typical clinical features in untreated diabetes mellitus are polydipsia, polyphagia, polyuria, and weight loss, but these occur only in the more severe forms of the disease. Because of the varied and nonspecific symptomatology of the disease, the actual diagnosis of diabetes rests upon laboratory evidence. It should be emphasized that all laboratory tests for diabetes are non-specific and abnormalities of one or all may occur in other diseases (e.g., hyperthyroidism, liver disease). However, the presence of other diseases, glycosuria and hyperglycemia are diagnostic.

- A Glycosuria. The presence of reducing substances identified as glucose in the urine is excellent presumptive evidence.
- B Hyperglycemia. The finding of an elevated fasting blood sugar and/or abnormal blood sugar level 2 hours after a meal containing 100 grams of carbohydrate or a dose of 100 grams of glucose is almost diagnostic in the absence of other diseases. This test should be performed, however, only after the patient has been on a high carbohydrate diet for at least 48 hours. It is well known that the previous diet influences carbohydrate tolerance. A high fat diet will decrease tolerance (even in nondiabetic type curves) and a high carbohydrate diet will increase tolerance.
- C Interpretation of Blood Sugar Tests. A normal fasting blood sugar does not rule out diabetes. If the 2-hour postprandial blood sugar level is over 140, one can be reasonably certain that the condition exists. If the blood sugar level is between 90 and 140, it is necessary to perform a glucose tolerance test to establish the diagnosis.

TREATMENT OF DIABETES MELLITUS

In order to treat patients with diabetes it is necessary that one thoroughly familiar with the following:

- 1 Insulin (probably of greatest importance)
- 2 Diet
- 3 Influence of exercise
- 4 Prompt treatment of complications
- 5 Newer agents effective against diabetes mellitus: Tolbutamide (Orinase®)

- 2 **Syring** In order to aid patients syringes are available calibrated in units (U) rather than cubic centimeters. Many of these syringes have 2 calibrations (U20 U40 or U40 U80) and it is important to have the patient thoroughly understand which scale he is using. It is advisable however to multiply a syringe having one calibration only. Special syringes are available for blind diabetic patients.
- 3 **Sites of injection** Insulin is usually administered subcutaneously. The site of injection is generally the anterior thigh but insulin may also be given in the lateral thigh in the arms or at the abdomen or in unusual circumstances subcutaneously in a other part of the body. It is important that the site be constantly changed so that the same site is not injected more often than once every 2-3 weeks. Crystalline and regular insulin may be administered I.V. to patients who have been taking insulin with out allergic reaction. *Do not give PZI NPH or lente insulin intravenously.*

Diet: Diabetic

The nutritional needs of the diabetic patient are not significantly different from those of normal individual. The principal question to be settled is the quantity and type of carbohydrate to be allowed in the diet. (For detailed food lists in making up diets see pp 46-53 and for examples of diabetic diets see p 57.)

Notes: Whenver possible diabetic diets should be made up in terms of household measures rather than weight for initially the extent a patient may gain weight by weighing is generally unnecessary.

The following factors must be taken into consideration in estimating the diet:

- A Caloric Needs (See p 47) The caloric needs of the diabetic are similar to those of non diabetic individual and the same variables must be considered. In general one should remember that the diabetic patient should be kept normal or slightly subnormal weight level and even permitted to become obese.
- B Protein Protein must be adequate and high protein diet are desirable because the available glucose (50%) from protein is relatively low for utilization than ingested carbohydrate. Allow 1-1.5 Gm. of protein per Kg. (0.45 Gm./lb.) of body weight should be given although 1 $\frac{1}{2}$ - 2 Gm./Kg. (0.7 - 0.9 Gm./lb.) is preferable.
- C Carbohydrate Carbohydrate should not be given in order to feed the patient. Protein should be given to 5% and 10% of total available carbohydrate to digest and absorb and allow available glucose to be obtained. The quantity of available carbohydrate in the diet is still unsettled. In general the view is taken that in the diabetic the aim is to keep the individual close to physiological normal as possible and hence to keep his carbohydrate at approximately normal level and to administer insulin if necessary to control a year. If hypoglycemia and glycosuria is general then 2-3 Gm. of carbohydrate per Kg. (0.9 - 1.4 Gm./lb.) of body weight is recommended with the aim of treatment if patient still increase with treatment gradually increase carbohydrate to 4 Gm. per Kg. (1.8 Gm./lb.) of body weight. This is a general rule and in some mild diabetes it

B Insulin Mixtures Intermediate insulin may be prepared by mixing a short acting or intermediate (commercial) and a long acting insulin (add last) in one syringe. This gives a balance between the immediate and the prolonged effects by modifying the mixture one can tailor the insulin requirements to individual needs. The mixtures usually employed are 2:1 and 3:1 (crystalline zinc PZI) or 2:1 and 3:1 (NPH PZI).

1 Points to remember in use of insulin mixtures

- Regular insulin must always be withdrawn into syringe before PZI (because of protamine excess in PZI)
- Same unitage of regular insulin and PZI must be used
- General effect of I/PZI mixtures: 1:1 largely PZI effect (little point to this mixture) 2:1 intermediate daytime/lighttime effect 3:1 greater daytime effect

2 Application of tailored insulin mixtures

- If glycosuria occurs in all fractional urines increase total insulin mixture
- If glycosuria occurs in fractional urines 1 and 2 only (daytime glycosuria) increase regular insulin mixture
- If glycosuria occurs in fractional urines 3 and 4 only (nighttime glycosuria) increase PZI in mixture

C Commercial insulin preparations come in various strengths (units/cc) in 5 and 10 cc ampules identified as follows

Potency Preparation	Color of Rubber Stopper	Color of Label
U20 Unmodified regular	Yellow	Yellow
U20 Crystalline	Yellow	Gray and blue or blue gray and yellow
U20 PZI	Not available	
U40 Unmodified regular	Red	Red
U40 Crystalline	Red	Red and gray
U40 PZI	Red	White label with red printing
U40 NPH	Red	Blue and white
U40 Globin zinc	Red	Red and brown
U40 Lente	Red	Gray label with red and black printing
U80 Unmodified regular	Green	Green
U80 Crystalline	Green	Green and gray
U80 PZI	Green	White label with green printing
U80 NPH	Green	Blue and white
U80 Globin zinc	Green	Green and brown
U80 Lente	Green	Gray label with green and black printing
U100 Unmodified regular	Orange	Orange

D Administration of Insulin

- Selection of insulin preparation. In view of the large number of insulin preparations available there is often great confusion regarding dosage. The effective it is necessary to place the patient on one type of insulin and have him become familiar with this type. One use and a limitation of subcutaneous that the volume per injection is kept to 0.25 to 0.5 cc. About 80% of patients are able to use U40 units.

B Compl ting Fa to s A larg number of factors adv ly aff ct the c rs f th patient with diabetes. All of th on dt s perate by altering the abs rption of glucos by int fer g with ca bohydr t oxidation by ausing excess v rb hyd t formation. The most imp t nt of these facto s c infections especi ly those of pyog na nature w th fev s d to emu. Any infectio s erio s in a d ab tic fo it ompl tely upa t th quilibrium established by th apy i wys i crea es the n d for i s hn nd is one of th mo t c mmon pre ipitating cau of k tosi d a ido is. The fo e ny nd all ind t ons in th diabet ca to b av ided wh ne e pos ble wh n th y o ur th y m st b t ted promptly a d v go o ly. Du ing sev re infect o s it ge er ally advi bl to di ont n PZI and to b gun th py in divid d do es 3 6 tim s daily with regula o crystall n in ln as n ed dt cove p tp dual glycos i.

C G ral F t P t ts with diabetes should hv n ly rmal hygi nic lives a p sible. Th y h ld b ass red of d quate st ho ld b abl to t t home if at all po sible and sho ld ngag in n o pation q i g at le st mod rat e is but m st av id st n ou o cupat ons of gr t tim p t n th y should hav good g n al kn wledge of diab tes.

STEPS IN THE MANAGEMENT OF THE DIABETIC PATIENT

Th r e may d q at m thod f m n gung di beti. Th follow g i plan u ed by th tho whi h f llt b pra t al d phys l g lly so d.

STEP A - DIAGNOSTIC WORK-UP

1. C mplet hi tory nd phy l xaminat fo diagno i and to l t th p f y o xting o mply ting d ea.
2. U naly fo qual t tive s ga n m ni g f ting u n p m n a d sp m ll t d 2 3 h aft a h m l H ga p t b k f a t and dia t a id.
3. Bl od s ga xamin t. Fast ga d 2 ho p tp dal l vels d t m n d o f say agl tol an e t t p f m d in ld ly patie ts in th p e of n al di. It advi bl t p f m a gluc t l and test with m lt in ga t det mine th appro m t n l th e h ld. If th i v y high (v 160 180 mg %) t may b n ess ry to e blood s ga l v l e a b k d qua y f th rapy rather than th glyc ria.

TL n (CALCULATION LID ARGUMENT OF DIST (e p 57 f xampl of d ab t d t)

1. D t m n th lo ic ds f th p t t. Thus i th m f th n n d a beti (p 47).
2. Cal ul t th p t in CHO and fat t t of th d t as outli d on pp 46 ff.

may be advisable to keep the carbohydrate level down to a minimum the use of insulin. However both for physiological and for psychological reasons the carbohydrate level should in no case be below 100 Gm per day.

- D Fat. After the carbohydrate and protein amounts have been determined fat is given to make up the remaining caloric requirements.
- E Vitamins. Patients with diabetes tend to develop vitamin deficiencies especially of the B complex. The reasons are not always clear but may be due to inadequate food intake, restricted diets, or increased requirements of improper utilization of vitamins. Deficiencies on adequate diets are rare if they occur treat as needed (see p. 59).
- F Frequency of Feeding. Diabetics should be given small frequent feeding rather than large meals. By frequent feedings the use of high protein intake and less concentrated carbohydrate food one can maintain a lower and more even blood sugar level with less glycosuria. An excellent plan is to divide the feedings into six meals: three regular meals and three small feedings (e.g. milk) at mid morning, mid afternoon and bed time.

Tolbutamide (N N D (O a))

An encouraging recent development in the treatment of diabetes mellitus is the introduction of a new oral tolbutamide (Orinase[®]) effective against certain forms of the disease. Tolbutamide is a hypoglycemic derivative although it has no antihyperlipidemic effect. It appears to stimulate the production of insulin by the beta cells of a pancreas which would otherwise produce the hormone (islet of Langerhans). It does not potentiate the action of insulin and is of no value in the pancreas of the elderly. Therefore tolbutamide is of no use (and should not be tried) in elderly diabetics (aged 60 and over) or in those diabetic patients who tend to develop ketosis. Its only use is in the older patient with a mild degree of diabetes which cannot be controlled by diet alone (i.e. mild to moderate nonketotic type).

Doage. Tolbutamide is applied in a total dose of 3 Gm daily divided into three doses and decreased rapidly to the minimal effective dose. The average maintenance dose is 0.5 to 1.5 Gm daily.

Toxic reaction is rare. Skin rash and gastrointestinal distress occur only occasionally. There have been no reported instances of granulocytopenia following use of tolbutamide. It may occur and should be borne in mind.

Other Factors Influencing Diabetes

A Exercise. Exercise enhances the oxidation of sugar and it diminishes the need for insulin. Therefore exercise in moderation is beneficial. However patients taking sulfonylureas should be cautioned against strenuous exercise without first notifying themselves previously with extra carbohydrate. (It is not uncommon to have a hypoglycemic reaction after a set of tennis.) When regulating a patient's hypoglycemic response approximately the same amount of exercise as will be required by his normal active life. This is true also of the patient with gestational diabetes.

time of glycosuria on the preceding day determines the re-adjustment to be made. The glycosuria at a given time must be kept at a minimum level. If the green reduction (or +) with the Zymatic test performed in any specimen is general, especially with the fasting urine, change should not be made frequently, simply because marked insulin action occurs only occasionally.

a. If all specimens are green, no adjustment of dosage or composition of insulin is necessary.

b. If glycosuria (greater than green reduction) occurs after breakfast after the noon meal, the regular insulin is increased.

If glycosuria (greater than green reduction) occurs in the afternoon after the evening meal, the regular insulin is decreased.

d. If glycosuria (greater than green reduction) occurs in all specimens, both regular and preprandial insulin are increased.

Amount of increase in insulin will vary with each patient. Generally, a very high glucose tolerance test of insulin follows.

(1) Yellow reduction (or ++). Add 5 to 10 units.

(2) Orange reduction (or +++). Add 10 to 15 units.

(3) Black reduction (or ++++). Add 15 to 20 units.

f. If the patient is on glycosuria (specimens remain blue) the patient should be observed for hypoglycemia and hemorrhoids should be examined. Adjustment of dosage may be made as indicated with the following.

4. Readjustment of the dosage of insulin. If variation of the insulin dosage and composition does not maintain the glycosuria at a minimum for a given period, the dietary intake for the preceding meal should be decreased and the intake for the next meal increased.

"P.D. - FOLLOW-UP OF PATIENT"

After patient has been adequately treated, he should be kept on a regular interval range of dosage to check for any change in the patient's individual status.

1. Hypoglycemia. Carefully question patient as to occurrence of any hypoglycemic attacks. If they occur, low insulin dosage during the time of day they take place.

2. Examine patient's urine. If all urine is entirely free of sugar, the patient is controlled (if renal threshold is normal). Beware of hypoglycemic reactions. How variable if all urine is blue in therapy. If patient tolerates new insulin properly and therapy. There is no indication to having some greater doses. (On the contrary, the risk is small that moderate glycosuria is not harmful if the metabolism of the body are being fulfilled. However, it is usually best to keep adequate control over patient's carbohydrate intake. (If the diet is entirely) If there is marked glycosuria in a given time, the insulin is adjusted accordingly.

3. Weight patient. Follow patient's weight to be sure that the weight is increasing and remaining stationary. A desired weight is the data accordingly.

- 3 Divide the diet into the following
 - a Three medium sized meals It is advisable to space the meals as far apart as possible (i.e. an early breakfast and a late dinner) This will spread the absorption of glucose over a longer period of the day
 - b Three small feedings to be taken between meals and at bedtime Milk and low CHO fruits are preferred for this

STEP C - DETERMINING THE INSULIN REQUIREMENTS

- 1 Determination of amount of glycosuria Have patient eat his diabetic diet for 1 day preferably without change in activity For the next 24 hours he is to collect and label fractional urines as follows (Patient voids just before breakfast and discards this specimen)
 - a Urine No. 1 All urine voided from breakfast to just before lunch This is pooled and a few drops taken for qualitative sugar The remainder is saved
 - b Urine No. 2 All urine from lunch to just before dinner Pool and save as above
 - c Urine No. 3 All urine from dinner to just before retiring Pool and save as above
 - d Urine No. 4 All urine from retiring to just before breakfast Pool and save as above

The few drops of each individual urine fraction are analyzed qualitatively for sugar and the remainder pooled for the daily total quantitative sugar
- 2 Calculation of approximate insulin requirements from quantitative urine sugar determinations Since roughly 1 unit of insulin will cover 2 Gm. of glucose the insulin needs in the uncomplicated diabetic can be calculated as follows

$$\frac{\text{Gm. of Glucose in 24 hour}}{\frac{\text{Urine Specimen}}{2}} = \text{Approximate No. of Units of Insulin Needed per 24 Hours}$$

The insulin (24 hour requirement) is generally given as NPH or as a mixture in a single dose $1\frac{1}{2}$ hour before breakfast The usual mixtures are 2:1 or 3:1 (crystalline zinc PZI) or NPH regular mixture

- a In severe or complicated diabetes because the patient needs insulin immediately these measures cannot be performed (see p. 403)
- b High renal threshold In certain elderly patients or those with renal disease who have a high renal threshold for sugar this method will be without value These patients must be controlled by the determination of the blood sugar levels while fasting and 1 hour after meals In these cases begin with small dose of long acting insulin (3-10 units/day) and increase as indicated by tests
- 3 Adjustment of insulin dosage and mixture The patient continues to collect his urine fractions as outlined above and the dosage and composition of the insulin mixture is determined each morning after completing the qualitative sugar analysis for the previous day Quantitative sugars are usually not necessary after the first day The amount and

- 2 Epinephrine (adrenaline) If patient is well nourished peacefully using short acting insulin and is not depleted of glycogen epinephrine 0.5 I.U. (8.15u) of 1000 oil solution but may cause reaction of vasoconstriction so that food may be taken by mouth.
- 3 Rectal feeding If patient is unconscious and I.V. glucose is not available (and if peripheral line is either not available or not feasible or unsuccessful) glucose by rectum may be given. Add 2 Tbsp. of syrup or honey to a pint of warm water and give slowly by rectum.

C Prevention of Relapse When patient is taking potassium zinc insulin development of complications should be carefully watched for danger of relapse. High protein foods such as milk should be given in addition to carbohydrate.

OTHER COMPLICATIONS OF INSULIN THERAPY

Allergic Reaction

Fortunately allergic reaction are very rare and most are localized. These individuals are generally sensitive to pork pancreas from which about 60% of commercial insulin made (the other 40% from beef). These patients should be given penicillin preparation (Special Insulin) which is put in 10 c ampules of U40. If patient is still sensitive to desensitization measures should be tried (p. 112).

Lipodystrophy

This is a complication consisting of atrophy of subcutaneous fat at the site of injection. This may be caused by improper rotation of injection sites but may also occur in spite of a regular therapy. These patients should use U80 or U100 insulin of the injection sites and maintain rotation at body sites which elicit little lipodystrophy.

COMPLICATIONS OF DIABETES

CHRONIC COMPLICATIONS

There is a tendency for these problems to develop with increasing frequency in diabetic patients than in nondiabetics and a few data indicate that are rather typically associated with diabetes. They are mentioned here to call attention to them. The therapy is generally that of adequate control of the diabetes and the therapy of the associated underlying disease. The most common diabetic complications are:

- A Arteriosclerosis Especially if the peripheral arteries of the legs. For therapy see p. 208.
- B Diabetic Polyneuropathy (See p. 358)
- C Diabetic Ocular Complications Included cataract which has increased frequency and retinopathy of which no form of therapy appears to be of any avail.

- 4 Draw blood for fasting blood sugar level to determine whether fasting hyperglycemia is being adequately controlled (This need not be done on every visit in fact it can be done quite infrequently once the patient is standardized)

COMPLICATIONS OF INSULIN THERAPY

HYPOGLYCEMIA (code No 574)

Hypoglycemia is the most common complication of insulin therapy and usually occurs when the diabetic fails to eat or engages in too strenuous exercise. It is manifested by weakness, hunger, irritability, faintness, and tremors and convulsions, all of which are relieved promptly by the administration of glucose. If a diabetic patient is seen unconscious and if diagnosis of coma or insulin reaction is impossible or in doubt, give 50% glucose I/V. This will definitely overcome the insulin reaction and will not generally harm the patient in diabetic acidosis.

Prophylaxis

- A Glucose Because of the danger of insulin reaction, the diabetic patient should carry several lumps of sugar or glucose lozenges at all times. If he feels the onset of a reaction, he should take some sugar.
- B Identification Card Every diabetic should carry a card with the following information:

I AM A DIABETIC AND TAKE INSULIN

If I am behaving peculiarly, give me sugar or hard candy or orange juice slowly. If I am unconscious, call an ambulance immediately, take me to a physician or a hospital, and notify my physician. I am not intoxicated.

My Name is _____

Address _____ Tel. phone _____

Physician's Name _____

Physician's Address _____ Tel. phone _____

Treatment

- A Mild Hypoglycemia If patient is conscious and able to swallow, sugar, glucose, or orange juice may be given.
- B Moderate to Severe Hypoglycemia Do not attempt to feed patient if unconscious. If patient is unconscious, one of the following methods may be used:
- 1 I/V glucose (treatment of choice) 20-50 cc (5-12 g) of 50% glucose I/V slowly. As soon as consciousness is restored, oral feeding may begin.

Diagn si

Diab ti ac d s is manif sted by heada he i itab lity d ow sin hyp rpn a and fever Naus vomiting diarrh a and abdominal pain may lso b pr s nt Th swe tish 'fruity' acc to breath m y b d tected On physical xamination the skin and muc s m mb nes re u ally dry blood p essu e low eyeballs soft nd pul su lly pid nd th dy

Principl of Th rapy

fo me ge y management ee b low

Th prn ple of th rapy wh th r th patie t p omato e o r in coma a e th sam It i imperative that a patie t in c do is be h pit l d nd t ated a medic l merg cy E h ca must be individuali ed

A I ulin n l g amounts i ne ary to b g about a r tu n to orm l metab l m *Use short-acting insulin never treat patients in coma with P.I. NPH or lente insulin* Th fi t dose f i insulin should be 100 300 unit on h lf should be giv n I V and the oth half ub utaneo ly Insulin may lso b added t I V fluids b g admi ste ed Be us of th mode of a tion of in lun (ee p 393) ther is no ne d t p at ner than in l 2 hours Th d se may the b r peated sub t o I V gi g 50 75 u ts ev ry l 2 ho s a need d until th keto u begi to d pp If ho k s p es t th in lun h ld be gi e I V beca e of the unrel ble b sorpt nd ung sho k of mate ial g ns b utaneo sly

B Glu ose In diabetic a idosi on s t ati g the k tos and a dos s a d n t the hype glyc mia and gly s ia Altho gh th pati nt with ido i may h v a high blo d s gar l v l the total availab e b hydrate st es m y actually b very low The for sin it s ne s ary t hav an adequate gl os pply pon whi h in lun n a t is overcoming cido is th se p t t sho ld be give gl co wh the blo d suga l vel has begun to f ll r pidly It has b sh wn that k tos can b re d c d by giving v ry la ge amou t of gl co to d i b tic

pat nts who s e dep ived of insulin The oon th norm l m t bolu pathways a e r established th soo r ess fat oxidati n c ses d k tonemia is over om In addit on it is po bl to p ipitat hyp gly mic e ction in a patient with l w s ga erv before th ket sis bro ght unde o t ol

C F u t se Inv rt S gar It ha be n shown that aft r I V infus o s f tos disappea from the blood st eam of diab tics s rap dly as it doe from o m ls It has be n sugg ted th r f re that this s gar be s b titut d f gl o e in th t at m nt of d b t i s be u it s ut l i ed in th bs e of insuli Howeve the is om videnc to show that in spite of its til t ion in the diabet it has no ant k i genl effect with t in uln Until this c t u l q t is ettl d o sh uld ont n e t us gl cose and in ulin i the manag m nt of diabeti c dosis

D Fl id and El ct lyt

- 1 Fluids m st be gi n t repla e th s lost by diuresis a d vomiting Th se ar u ally best given I V
- 2 Ad q t odium hloride is v ry importa t This repla s fixed bas in th xt e ellular fluid a d so h lps in ove oming th a ido is As a re ult of ketosis the los of

- D Renal Complication Inter-capillary glomerulosclerosis characterized by hypertension albuminuria and edema. Treat as for glomerulonephritis (see p 203)

ACUTE COMPLICATIONS OF DIABETES

When the amount of insulin in the body is inadequate for metabolic needs abnormal metabolism results with ketone body formation and finally with acidosis. Infection which causes an increased demand for insulin usually precipitates ketosis. There is an early or mild phase and a late or severe one.

- A Diabetic ketosis Without Acidosis CO_2 content or combining power is normal or slightly depressed (above 50-60 Vol % or 27 mEq)
- B Diabetic Acidosis Reduction of CO_2 content or combining power (below 50 Vol % or 27 mEq). The patient may be conscious pre-comatose or comatose.

DIABETIC KETOSIS (Without Acidosis) (code No 543)

In this disorder ketone bodies are found in the urine and their presence establishes the diagnosis. Examine the patient for infection or other precipitating factor. The fluid and electrolyte balances are undisturbed.

Treatment

Patient should be hospitalized for regulation if ketosis is severe.

- A Treat any infection which may aggravate the disorder and metabolism.
- B Diet Arrange diet to contain 3 equal feedings with interval feedings between each meal and in the evening.
- C Insulin
- 1 If ketosis is very severe use only short-acting insulin. Give insulin to cover each meal as necessary until the urine is free from ketone bodies. Then begin reducing insulin dosage slowly as tolerance to carbohydrate improves.
 - 2 If ketosis is not severe treat and regulate as in complicated diabetes.
- D Follow up When ketonuria has cleared patient is managed as for uncomplicated diabetes according to the severity of his disease (see p 399).

DIABETIC ACIDOSIS (Diabetic Coma) (code No 542)

When ketone formation is proceeding at a rapid rate the fluid and electrolyte balance and pH of the body are upset (see below). The ketone bodies are organic acids which replace the HCO_3^- in the body and also are excreted from the body combined with fixed bases. The loss of fixed base and the disturbance of the buffering systems leads to acidosis. The increase in the glucose in the blood produces decreased osmotic pressure of the body fluids.

insulin per gram of sugar (5-10 units insulin per liter) and 20 mEq potassium and possibly phosphate. The urine should contain sugar at all times to avoid hypoglycemic reactions.

- b As soon as reports come from laboratory if CO_2 combining power is below 5 mEq/liter (10 Vol %), calculate amount of sodium lactate or sodium bicarbonate desired (see p. 13) and administer immediately. (To administer sodium bicarbonate 1 V. multiply desired chemically pure sodium bicarbonate in 100-300 cc cool distilled water and administer. Do not heat or sterilize the solution.)
- c Gastric lavage may be performed with introduction of 200 cc of physiological saline or 5% NaHCO_3 .
- d As long as patient is unconscious administer 5% glucose in saline or other salt solution as indicated (about 60 drops per minute). See p. 405.
- e As soon as patient is conscious and able to swallow give fruit juice (200 cc orange juice with 1 tablespoon honey syrup or glucose) every 3-4 hours until ketonuria has stopped. Stop 1 V glucose and fluids.

B Follow up

1 Potassium deficiency. After 4 to 8 hours of administration of 1 V fluids watch patient carefully for potassium deficiency (increased weakness, respiratory distress) and check the ECG (see p. 14). Give oral potassium in long position (see p. 21) as indicated. If not yet able to begin administration of potassium as soon as the comatose state is begun but still not settled. When patient is able to swallow give complementary potassium salts by mouth as this is the safest route.

2 Oral feedings and fluid. If ketonuria is disappearing or is rapidly improving (usually in 24-48 hours) and the patient is conscious the following may be given:

- a Small frequent feedings of liquid and semi liquid foods containing 50-75 Gm glucose and protein (as milk) every 3-4 hours day and night and cover with 5-35 units regular insulin every 4 hours.
- b Food fluids by mouth.

Examine urine for sugar and ketone bodies every 3-4 hrs. Regular diet. After 24-48 hours if patient shows steady improvement place on regular diabetic diet and begin regulation as outlined on p. 393.

DIABETES ASSOCIATED WITH OTHER CONDITIONS

PREGNANCY

The management of the pregnant diabetic is little different from that of a young diabetic.

- A During the early period of pregnancy there is often a lowering of the normal threshold and considerable lability of the blood sugar level.

sodium chloride from the body may be as great as 30 Gm (50% of average total body sodium) in 24-48 hours. In the mild case sodium chloride needs to be replaced and sodium chloride solution with glucose is usually adequate fluid therapy.

- 3 Replacement of bicarbonate buffer. As the ketone bodies are excreted or oxidized CO_2 is formed which replaces the disappearing ketones and the CO_2 -combining power returns to normal. However in patients with severe uncomplicated metabolic acidosis it may be advisable to administer more rapidly available HCO_3^- and fixed base (i.e. Na^+). This may be given I.V. as sodium bicarbonate or M/6 sodium lactate.
- 4 Potassium replacement. As sodium is administered (as sodium chloride, sodium bicarbonate or sodium lactate) and glucose is metabolized and stored the potassium which has entered the extracellular fluid migrates rapidly intracellularly or is washed out with the fluid through the kidneys. When this occurs there may be a temporary and dangerous extracellular potassium deficiency with weakness, respiratory distress and at times cardiac arrest. Solutions containing potassium must be given to correct this and generally when I.V. glucose becomes indicated potassium may be added to the infusion mixture (see p. 21). It must be used with extreme caution in the absence of adequate urinary output. The level may roughly be checked with the ECG (p. 14).

Treatment.

A. Emergency Measures. The following is an outline of the steps that may be employed in the average patient in diabetic coma; however each case must be individualized and therapy modified as necessary according to the needs of the patient.

- 1 Hospitalize patient. Keep patient warm, avoid excessive warmth. Avoid the use of barbiturates and narcotics.
- 2 If in SHOCK treat with I.V. plasma and other shock measures especially vasopressors (see p. 27).
- 3 Blood chemistry. Draw blood for CO_2 -combining power and blood sugar also for sodium, potassium and chloride if these tests can be performed.
- 4 Give insulin at once.
 - a Through same needle as for drawing blood give 30-100 units of regular or crystalline insulin I.V. immediately as well as a like amount subcutaneously.
 - b Repeat insulin giving 50-75 units subcut every 1-2 hours until there is rapid diminution in blood or urinary sugar.
- 5 Catheterize patient. An indwelling catheter may be left in place allow this to drain continuously. Examine spot (per os) urine specimen very hourly for ketone bodies and sugar.
- 6 Fluids, electrolytes and glucose.
 - a Begin I.V. infusion of saline. May also begin lysis of saline M/6 sodium lactate or other indicated solutions at same time (see p. 15). As soon as urinary sugar has changed to dilute or negative then change I.V. fluids to 5% glucose in saline to which 1 unit/l.

B F T um Although in reas d bohyd ate tolera de
 velop rap dly a a sult of t um th p inciple dange in a
 t t d d b ti who in j red s th possib lity of having a
 se e hypogly mi react on beca h f ile to eat Th e
 fo if th pat e t onscous give sw t ned ra ge ju
 or ca dy by mo th if s g ry is e ry giv 5% gl co e
 l V in wat o sal lowly One m y add l unit insulin per
 2 3 Gm gl cos t th nf so h w ve th need not so
 m h fo in lin as f gl co to avo d hypogly ma When
 s g ry h s b n compl t d t at acco ding to verity of
 dise s (ep 394)

Elect v S gery

A t t l H p t l M a

- 1 P tent ho ident ho p t l s veral d ys befo e surg ry
- 2 D ti e p ot m n inc in ulin
- 3 Th d ab tes sho ld be br ght nd opt m m co t of with
 gul r or c y tall n ins lin
- 4 There should be o keto :

B D ing nd Aft S g ry

- 1 N f o d o in ulin sh uld be dm nist d on the m rning of
 gery
- 2 Ma gem nt during s gery
 If th p tent diab t is mild nd has b en p operly
 o t l d if h d es n t tend to de l p k tos s and if
 th rgery i ot t ext n live h m y be op rated on
 w thout f o d o i sub
 b If th p t t diab t s a mode at or ve o if e
 te s a rge y m t b p r f m d begin info n of 5%
 gl os in lin o wat to wh h has be add d i unit
 gula r ry tall n i sul pe 2 Gm gl o C n
 ti u inf s on th o gh t su gical p ocedu e Giv inf
 sion i abo t 60 70 d ops per mi t
- 3 Afte s gery patie t sho uld hav small freq t f dings
 (50 75 Gm a bohyd te) ve y 3 4 hou cov red w th
 15 25 nit of ryst lin i ulin sub t u ly b fo e th
 m l These small fe dings ont u d ntl mal
 nut ition s be stablsh d
- 4 If ga t ont stinal s gery has b n p r f m d nd pat nt
 cannot take food by mo th n trit a can b t be m unta ed
 by pa nt l m th d giv 1000 c 5% gl co in 5"
 muno e d s l t i n l V slowly ov a per od f 4 ho r
 This h ld b ove d w th 15-40 unts of rystalln in ulin
 bc t b fo beginning inf ion Th lts s per day s
 s r g q : m nt This the apy may b ondu ed
 nt l r l utrition can be es med

HYPERINSULINISM

(Adenoma code No 871 8044A)

(Without Tumor code No 871 784)

Hype i ulinism a d by exc s v p odu t on of in ulin
 a d manif ted by att ke of w aka hungre ir it bility faint
 n a d t eme a d co v l ions all f whi h u ally occ o

- B During the latter three months there is often a marked decrease in glucose tolerance necessitating increased insulin dosage. This is not universal however and many go through pregnancy without significant changes in tolerance.
- C Before the onset of labor and delivery it is advisable to change to short acting insulins to avoid possible reaction from lack of food.
- D In view of work suggesting sex hormonal imbalances in pregnant diabetics therapy with estrogens or progestone or both has been advocated as being of value in diminishing fetal mortality. However carefully controlled studies using modern diabetic treatment methods show as good or better results without resorting to this expensive and troublesome procedure.
- E Since many diabetic pregnancies go beyond expected term or the infants are unusually large it has been suggested by many that pregnancy be terminated at about 36 weeks. The preferred method appears to be cesarean section.

The Care of the Infant

Treat all infants of diabetic mothers as prematures. Keep infant in incubator under oxygen for first several days. Observe the newly born infant carefully for the first 72 hours for hypoglycemic reactions that may occur supposedly as a result of islet cell hyperplasia. This is more apt to occur in the newborn of poorly controlled diabetics.

SURGERY

Surgery in the diabetic at present presents little hazard over that of the surgical procedure per se. However there are certain problems peculiar to the diabetic and these problems naturally vary with the severity of the disease and the urgency of surgery.

Emergency Surgery

- A. F. Non Traumatic Conditions Usually diabetics requiring emergency surgery for non traumatic disorders are in ketosis with or without acidosis and require immediate treatment of their diabetes. These patients should be treated as patients with acidosis or coma (the latter if a general anesthesia is to be used). The general program should be as follows:
- 1 Draw blood for CO₂ combining power and blood sugar also for electrolyte panel (sodium potassium chloride) if possible.
 - 2 Begin 5% glucose in saline infusion I.V. slowly (not over 70 drops per minute) and continue infusion throughout surgical procedure. One unit of insulin per 2 Gm glucose may be added to the infusion (25 units for each 1000 cc of 5% glucose).
 - 3 Give 50 units short acting insulin I.V. if ketosis is present.
 - 4 After returning from surgery continue therapy as for diabetic coma (see p 404) until oral feeding can begin and ketosis and hyperglycemia are controlled.

Chapter 16

HORMONES AND HORMONE LIKE AGENTS

PITUITARY AND PITUITARY LIKE HORMONES

The pituitary consists of two parts

- A Posterior which is part of the CNS
B Anterior portion which is devoid of direct innervation
Certain pituitary like hormones that influence the gonads are laboured by the placenta during pregnancy

ANTERIOR PITUITARY HORMONES

All of the anterior pituitary hormones are protein substances and must therefore be administered parenterally to be effective if taken by mouth. They are digested by the digestive enzymes. In general with the exception of the growth and lactogenic hormone which has effects as a direct stimulant to the growth of the anterior pituitary hormones appear to have a regulatory function rather than gland of internal secretion.

In the last few years several of these hormones have been prepared in pure form to permit detailed study of the growth, adrenocortical (ACTH), lactogenic, thyroid stimulating (TSH), follicle stimulating (FSH) and interstitial cell stimulating (lutinizing) hormones. There may be others in the anterior pituitary but they have not yet been fully identified. Of the preparations only ACTH and thyrotrophin are at present commercially available.

Corticotrophin (ACTH)

Corticotrophin has been shown to have remarkable effects in increasing many of the processes which are not satisfactorily influenced by other therapeutic agents. Its effects are entirely mediated by the stimulation of the adrenal cortex. Corticotrophin is a protein of molecular weight 4500 and contains 191 amino acids. It has been found to have similar and a marked physiological effect as the hormone itself.

A Metabolic Effect in Human ACTH in doses of 40 units in normal human being produces the following metabolic effects:

- 1 Increase in fluid and electrolyte balance and phosphate
- 2 Retention of sodium and electrolyte retention of water
- 3 Elevation of fasting blood sugar and diabetic glucose tolerance test
- 4 Increase in excretion of uric acid

410 Hyperinsulinism

an empty stomach long after meals and are relieved promptly by the administration of glucose. Hypoglycemia noted during episodes usually is below 40 mg %. A glucose tolerance curve drops to exceedingly low levels often only after 5-6 hours is characteristic. Hypoglycemia of hepatic, nervous or other endocrine gland origin must be considered in establishing the diagnosis.

Treatment

A Emergency Treatment As for hypoglycemic reaction from insulin overdosage (see p. 402)

B General Measures

1. **Corticotropin (ACTH) or the cortisones** The administration of these drugs (for their hyperglycemic effect) has been shown to be of considerable benefit in the management of some children suffering from this condition. Some children without adenomata have been successfully maintained or treated intermittently with these drugs.

2. **Diet** High protein, high caloric, high fat, low carbohydrate

a. The diet is low in carbohydrates in order to avoid stimulation of the pancreas to elaborate insulin. Rapidly utilized carbohydrates are replaced by slow acting ones (e.g. 5-10% vegetables and fruits and bananas and apples). Protein is an important source of slowly liberating carbohydrate which apparently has less stimulating effect on the pancreas and is useful to supply added calories.

b. **Small feedings** The diet is best divided into six or more meals a day. It may be necessary to feed the patient at regular intervals throughout the entire 24 hours. If the hypoglycemia is severe as this it is advisable not to prolong medical therapy but to prepare the patient properly for surgery.

3. **Sedation** Phenobarbital, phenobarbital 15-30 mg (1/4-1/2 gr) q.d. may be valuable in reducing nervous irritability.

4. **Restriction of physical activity** Exercise increases utilization of glucose thereby exaggerating the effect of excess insulin. If exercise is unavoidable, sedation should be preceded by supplementary carbohydrates.

5. **Identification** and Patient should carry a bracelet or card similar to that used by a diabetic (see p. 402).

6. **Emergency CHO** Patient should be required to carry a small supply of rapidly available carbohydrate (candy lumps of sugar) at all times. He is to avoid taking these except when definitely indicated.

C Surgery Complete excision of hyperplastic adenoma or islet tissue when this is found to be the cause.

Chapter 16

HORMONES AND HORMONE LIKE AGENTS

PITUITARY AND PITUITARY LIKE HORMONES

The pituitary consists of two parts

A Posterior part which is a part of the C N S

B Anterior part on which is the old of diet u al t m v t

Certain pituitary like hormones that influence the growth of the
laboratory by the placental growth

ANTERIOR PITUITARY HORMONES

All of the anterior pituitary hormones are secreted in substance
and must therefore be administered parenterally to be effective if
taken by mouth they are digested by the digestive enzymes In
general with the exception of the growth and lactogenic hormone
whose effects are not mediated directly through the gland the
anterior pituitary hormones appear to have a regulatory function on
the glands of the endocrine system

In the last few years several of these hormones have been pre-
pared in pure form These include the growth and corticotropic
(ACTH) luteinizing thyroid stimulating (TSH) follicle stimulating
(FSH) and interstitial cell-stimulating (luteinizing) hormones
The last may be either for the anterior pituitary but they have
not been fully defined Of the pituitary hormones only ACTH
and thyrotropin are important clinically

Corticotropin (ACTH)

Corticotropin has been shown to have remarkable effects in
regarding many diseases which are not at first obviously in-
fluenced by other therapeutic agents Its effects are entirely mediated
by the stimulation of the adrenal cortex Corticotropin is a part of
of the molecular weight and is a peptide derived from the
beef and human and a marked physiological effects as
the hormone itself

- A Metabolic Effects of H m s ACTH in adequate doses in nor-
mal human beings produces the following metabolic effects
- 1 Increase in the excretion of sodium potassium and phosphorus
 - 2 Retention of sodium and secondary retention of water
 - 3 Elevation of blood sugar and dibutyryl glucocorticoid
and uric acid
 - 4 Increase in the excretion of uric acid

- 5 Increased urinary 17 ketosteroids and cortiosteroids
- 6 Fall in circulating eosinophils and lymphocytes and leucocytosis of polymorphonuclear leucocytes

B For clinical effects uses and dosages see page 423

Pituitary Growth Hormone (PGH)

Purified PGH has been employed in normal humans, pituitary dwarfs and panhypopituitary individuals. In no case has there been any evidence of growth as measured either by metabolic effect or actual physical growth. The older crude growth hormone preparations have likewise been of no benefit under controlled experimental conditions.

Lactogenic Hormone

Has not been employed in human research. Its presence is necessary for the initiation and apparently for the continuation of lactation in breasts which have been prepared for lactation by estrogens and progestins during pregnancy.

Follicle Stimulating Hormone (FSH)

FSH has different actions in male and female. In the female FSH stimulates the development of ovarian follicles. In the male it stimulates the germinal epithelium of the testis to produce spermatozoa. It apparently has no effect on the Leydig cells, hence does not influence testosterone secretion. Pure FSH has not been used clinically but in cases of hypogonadotropic eucholesterolemia a purified preparation has been employed after initial stimulation of Leydig cells with chorionic gonadotropin to initiate spermatogenesis (see page 386). At present no good FSH preparation is commercially available.

Interstitial Cell-Stimulating Hormone (ICSH) (Luteinizing Hormone)

A In the female ICSH apparently has a dual action:

- 1 Stimulates growth of theca cells
- 2 Transforms the mature follicles into corpora lutea

B In the male it stimulates the Leydig cells of the testis with resultant testosterone secretion.

There is no good commercial pituitary ICSH. Clinically ICSH is substituted for by use of chorionic gonadotropins which have a similar action (see page 413).

Thyroid Stimulating Hormone (TSH)

TSH is exceedingly effective in stimulating the thyroid gland. It has limited clinical usefulness at present. Its principal uses are to differentiate pituitary hypothyroidism from primary hypothyroidism. It has also been used in an attempt to stimulate metastatic thyroid cancer to take up additional iodine for therapeutic purposes.

Recently it has been advocated for the treatment of thyroiditis but its place in the management of this disease is still open to question.

Dosage: 5-10 U.S.P. units I.M. every 12 or 24 hours for 2 days. Repeat ^{131}I uptake or protein-bound iodine. If uptake or PBI is increased primary hypothyroidism is not present.

Other Hormones

The pituitary probably exerts its effects at several points (see page 412) but their exact physiological roles are not yet known.

POSTERIOR PITUITARY HORMONES

The posterior pituitary hormones are polypeptides composed of 8 amino acids. Their exact chemical structures have been determined and they have recently been synthesized. Like the anterior pituitary hormones they are effective only when administered parenterally (i.e. i.m.). They exert their action

1. They raise blood pressure (pressor action) (an anesthetized animal)
2. Cause fluid retention without osmotically equivalent sodium retention (antidiuretic action)
3. Cause uterine contractions (oxytocic action)

To date there has not been a separation of the antidiuretic from the pressor principle. They may be identical. The oxytocic factor may likewise have some pressor effect.

Clinical Uses

- A. Antidiuretic - Pressor principle is used primarily for the treatment of diabetes insipidus also to prevent and treat abdominal distention (F. D. bet. Insipidum pag. 366)
- B. Oxytocic - is employed in obstetrics when indications for the induction of uterine contraction exist.

Preparations Available

Name	Action	How Supplied	Average Dose
Vasopressin Tannate NND (Pit. in Tannate)	Antidiuretic pressor	Olysolton 5 units/c	0.3 i.u. i.m. (5-16 u.) q 1-2 hr.
Vasopressin Injection USP		Aqueous solution 20 units/	0.25-0.5 c (4-8 u.) q 3-4 h. s.i.m.
Oxytocin Injection USP (Pit. in)	Oxytocic	Soluton 10 units/c	0.3 i.u. i.m. (5-16 u.) as indicated

PITUITARY LIKE HORMONES ELABORATED BY PLACENTA

The most important of the pituitary like hormones is that elaborated by the placenta during pregnancy. The hormone is referred to as human chorionic gonadotropin. Its physiological action is almost identical with that of ICSH. It has been shown that this hormone apparently functions only if an intact anterior pituitary gland is present. It is of no value in inducing spermatogenesis or ovulation or maintaining corpus luteum by itself. Many of its alleged effects have been due to the presence of FSH whose action the presence of chorionic gonadotropin may potentiate.

Clinical Indications

A. In the Male

1. Cryptorchidism - In a few select cases chorionic gonadotropin may induce descent of testis.
2. Hypogonadism - Chorionic gonadotropin is useful in some

types of hypogonadism although testosterone medication is generally preferred

- B In the Female Chorionic gonadotropin may aid in inducing ovulation and maintaining corpus luteum in a few selected cases of sterility if adequate FSH is present

Preparations Available

- A Chorionic Gonadotropin N N D derived from the urine of pregnant women is available commercially under a wide variety of trade names. It is marketed in ampules of 100 500 1000 and 3000 I U per cc.
- B Equine gonadotropins derived from the serum of pregnant mares is also available commercially. This is a mixture of FSH and LSH. It is not generally recommended because of its marked sensitizing effect and production of antihormones by protracted use. Only short courses should be employed.

Average Dosage

Usual dose: a range from 200-1000 units every day or every other day

THYROID HORMONE

The simple principle of the thyroid gland appears to be the iodine-containing amino acid thyroxine. Thyroxine probably never occurs in the free state in the organism but is contained in a protein molecule thyroglobulin. Another iodine-containing amino acid diiodotyrosine with weak physiological effects is also found in the gland. Triiodothyronine (or monothyronine Cyt m l¹⁰) has been isolated from the thyroid. It is about 4 times as potent as thyroxine and acts more rapidly. Its exact physiological role is unknown. The action of the thyroid hormone is that of a general cellular metabolic stimulant with respect to basal oxygen consumption (see increased metabolic rate). Its exact mode of action is unknown.

Method of Administration

Thyroid hormone either in the form of thyroglobulin (desiccated thyroid) thyroxine or triiodothyronine is effective when taken orally. There is a marked difference in the rate of metabolic responses between triiodothyronine and thyroxine. In the case of thyroxine little effect is noted after a single dose for about 24 hours and the maximal effect is not reached for 8-10 days. After the medication is stopped there is a slow loss of the effect depending on the initial B.M.R. and the level reached during thyroid medication. In general it may be stated that at least 4-6 weeks must elapse after thyroid medication has been discontinued before one can be reasonably certain that the effect has been dissipated. In the case of triiodothyronine the peak effect is reached in 12-24 hours and the effect is over in about 9-14 days.

Contraindications

Thyroid hormone is indicated only in thyroid deficiency states. Its use as a general metabolic stimulant is not indicated and is worthless. It has been shown that patients with thyroid deficiency rarely

require over 0.13 Gm (2 gr) of Thyroid U.S.P. daily. Patients with uterine fibroids can easily tolerate 0.3 to 0.5 Gm (5 to 7½ gr) monthly without any effect on B.M.R. although the 131 I uptake is suppressed. A good general rule is that if a patient requires over 3 gr of Thyroid U.S.P. daily his need for thyroid medication should be questioned.

Preparations Available

- A Thyroid U.S.P. B.P. (desiccated thyroid). This is the preparation of choice. The evidence indicates that any of the commercial preparations on the market containing more or less iodine than the official dose are any less toxic. To avoid confusion in dosage always use the official thyroid.
Dosage 0.065 to 0.2 Gm (1.2 gr) daily.
- B Thyroid Thyroid Tablets. There is no advantage of the crystalline thyroid over desiccated thyroid. It is approximately 300 times as potent as thyroid and small changes in dosage may lead to illness.
- C Sodium L-thyroxine N.N.D. (Tetraiodothyrene Cytom 1®). Meripol tablets disappear from the system with a half-life of 3.4 times longer than thyroid. Excrete metabolites without elevating protein-bound iodine. Hypothyroidism. Average maintenance dose 0.05 to 0.15 mg daily.

PARATHYROID HORMONE

Parathyroid hormone is a protein substance derived from the parathyroid gland. It can also be employed clinically for more than a few days, week, for the production of an increase in calcium (probably through production of antihypocalcemic agents).

Parathyroid hormone has many effects on calcium metabolism. It affects the release of calcium from bone stores. It affects the release of calcium from the diet. It affects the release of calcium from the diet. It affects the release of calcium from the diet.

Because of the fact that development of parathyroid hormone is two other preparations are employed in its place. They are dihydroxycholesterol (A.T. 10) and vitamin D. Both of these are tablets and are effective by mouth. Although the first A.T. 10 was the preparation of choice, the new preparation of vitamin D is almost equally effective.

Classification

Parathyroid hormone is indicated only in a hypoparathyroidism (functional removal of parathyroid gland).

Preparations Available

- A Parathyroid Injection U.S.P. Aqueous solution contains 100 units per cc.
Average dose 50-100 units (0.5-1.0) 3 to 5 times daily.

I M as indicated

B Dihydrotyrosol N N D (Hytakerol®) Each cc contains
125 mg of dihydrotyrosol in oily solution
Dosage See page 378

C Calciferol U S P B P (Vitamin D₂) 15 mg (160 1/2 gr)
daily (has a potency of 40 000 units per mg) (see page 378)

THE ADRENAL HORMONES

ADRENAL CORTEX

The hormones of the adrenal cortex are all steroidal substances. To date over 30 different steroids have been isolated and identified from animal adrenal glands or adrenal venous blood. The vast majority of these steroids have no demonstrable metabolic effect.

Questions have been raised as to whether or not all the steroids isolated from the adrenal cortex are truly naturally occurring or whether they are artifacts produced in the chemical laboratory. Recent isolation of hormone from blood obtained by catheterization of renal veins shows that most of the hormone (about 90%) is 11-hydroxycorticosterone (Compound F) and about 10% corticosterone (Compound B). In general it may be stated that the best demonstration of the effects of adrenal cortical hormone is that seen following administration of ACTH (see page 423). Recently aldosterone has been isolated from adrenal. This hormone appears to have only sodium and water retention and potassium losing effect. It is about 20 times as potent as desoxy corticosterone (D O C A). Hormones with estrogenic and androgenic effects have also been isolated.

Clinical Preparation

Of the dozen steroids isolated, three have had significant clinical use and trial. In addition to these two important chemical derivatives have been prepared.

A Desoxycorticosterone Acetate U S P Deoxycortone Acetate B P (D O C A) It is only a significant metabolic effect sodium and water retention and a decrease in urinary potassium excretion. In this respect it is approximately twenty times as potent as cortisone. It has no effect on carbohydrate or protein metabolism.

B Cortisone Acetate U S P B P

- 1 The principal metabolic effect of cortisone
 - a Retention of some sodium and water
 - b Increased excretion of nitrogen, potassium and phosphorus
 - c Increased blood sugar and ability to maintain blood sugar levels during fasting in Addisonian patients
 - d Return of EEG pattern to normal in Addisonian patients
 - e One of the most important effects is the adrenal cortical atrophy which results with prolonged use. This is probably due to excessive ACTH inhibition. The result is the absence of the normal response of the pituitary to stress.

- 2 For clinical effect, a dose of

C Hydrocortisone U.S.P. This compound has recently been
 established as a local (gingival) use
 It closely resembles that of cortisone. It is probably
 about 1 1/2 times as potent as cortisone on a weight basis. The
 molecular formula of hydrocortisone appears to be identical with
 those of cortisone.

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5 times a g e t as th t of D O C A and is flim t d
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E P d o N N D P d n i o l o N N D a d M thyl
p d n i (M d iⁿ) P d s o e a n d p d n o l o e a the
d h y d o i 2 d r t of c o t o a d h y d r o o r t e
p e t l y Ther app e r t o b l t l e i f a n y q t t a t e d f f
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t h t h y (1) s e l t l e i f a y s d i m r t e t o n (2) p s s b l y
u s l e s s p t s i m l and (3) a b o u t 3 5 t i m e s p o
t i t i n t h e r m t b l h n d t h e r p t i c a t i o F o t h e s
a s t h y r g n l l y r p l c i g t h e i r p t o m p o d s
w h e r t h e s h o m o e s a u s d f o o t h t h a p l a m e t
t h a p y (e g d e n l i n s f f c e c y) M t h y l p e d n i s o l i s
b t l 1/2 t i m e s p o t t s t h e p r e d n i s o e a n d p e d i s o l o e
I t p o t o h o t h a d i g

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m ly mpound F (b t 2 mg / c) w th les mpound E

P p t t A v l b l

A D y t t e A e t t U S P D y c o r t o n e A t t
B P (D O C A) o r D e y t o s t r o T m t h y l t t
U d o l y f o r p p l m t a r y m t a n o f A d d s d i s
1 B l t b l e t s 2 m g t b l e t s f r a b r p t n f o r m m u u s
m e m b e D O C A i i f f e t i v e w h n e w f l o w e d
D s g 1 2 2 t a b l t s d a i l y d i o l d i n t h b a l g u t t r I t
l m t q l l y f f t i e u n a g i v d a s w h n i n j c t e d
2 S o l t i n u n s m i l 5 m g (1 / 1 2 g r) p
D a g e 1 3 m g (1 / 2 0 1 / 2 0 g r) I M d i l y f m a i n t n a n e
3 P l l t s U S P 7 5 m g (1 1 / 4 g r) o 1 2 5 m g (g r) a c h
f o r u b c t i m p l a n t a t
D o g O 7 5 m g p e l l t f e a c h m g o f D O C A e
q u d b y i n j t i o p t o 3 m g (1 / 2 0 g r) p d y I f e

- 1 M s s i d e t e d
- B D h d r o t a c h s t e r o \ \ D (H i k e r o l[®]) Each cc contains
1 25 mg of d h d r o t a c h s t e r o i n o i l y s o l u t i o n
Dosage See page 378
- C Cal i f e r o l L S P B P (Vitamin D₂) 1 5 mg (1 63 1/2 gr)
daily (na s p o t e n c y o f 4 0 0 0 u n i t p e r m g) (see pag 3 8)

THE ADRENAL HORMONES

ADRENAL CORTEX

The hormones of the adrenal cortex are all steroidal substances. To date over 30 different steroids have been isolated and identified from animal adrenal glands or adrenal venous blood. The vast majority of these steroids have no demonstrable metabolic effect.

Questions have been raised as to whether or not all the steroids isolated from the adrenal cortex are truly naturally occurring or whether they are artefacts produced in the chemical laboratory. Recent isolation of hormones from blood obtained by catheterization of renal veins shows that most of the hormone (about 90%) is 11-hydroxycorticosterone (Compound F) and about 10% is cortisone (Compound B). In general it may be stated that the best demonstration of the effects of adrenal cortical hormone is the demonstration of the effect of corticotropin (ACTH) administration (see page 422). Recently aldosterone has been isolated from adrenals. This hormone appears to have only sodium and water retaining and potassium losing effect. It is about 20 times as potent as desoxycorticosterone (D O C A). Hormones with estrogenic and androgenic effects have also been isolated.

Clinical Preparations

Of the adrenal steroids isolated, there have been two significant clinical uses and two. In addition to these two important chemical derivatives have been prepared.

- A De o x y c o r t i c o s t e r o n A e t a t e L S P Deoxycorticosterone A etate
B P (D O C A) Its only significant metabolic effect is on
sodium and water retention and increased urinary potassium ex-
cretion. In this respect it is approximately twenty times as
potent as cortisone. It has no effect on carbohydrate or protein
metabolism.

- B C o r t i s A c t i v e U S P B P

- i The physiological effect of cortisone are
 - a Retention of sodium and water
 - b Increased retention of strontium, potassium and phosphorus
 - c Increased blood sugar and ability to maintain blood sugar level during fasting in Addisonian patients
 - d Return of EEG pattern to normal in Addisonian patient
 - e One of the most important effects is the adrenal cortical atrophy which results with prolonged use. This is probably due to endogenous ACTH inhibition and may result in the absence of the normal response of the pituitary-adrenal axis to stress.

2 For clinical effect and use see

4 Rare uses in cardiology e.g. Stokes Adma syndrome and cardiac arrest

5 Diagnostic test of hepatic glycogen storage

B Preparation Available

- 1 Epinephrine Injection U.S.P. Adrenaline B.P. 1 mg / cc (1:1000) Administered generally subcut may be given I.M. and even I.V. If diluted in 1 liter of solution Do age 0.2-1 cc (3-15 μ) as indicated
- 2 Epinephrine Inhalation U.S.P. 10 mg / cc (1:100) For inhalation only
- 3 Epinephrine in Oil Injection U.S.P. 2 mg / cc (1:500) Administered only I.M. Usual dose 0.2-1 cc (3-15 μ)

Actions of (Norepinephrine)

A Clinical Uses Adrenaline is used almost exclusively for its vasoconstrictor effect. Used in a wide hypotensive states (surgical and non surgical shock) central vasomotor depression and hemorrhage (see p. 27)

B Preparation Available Levartre 1 Bitartrate U.S.P. (Levophed[®]) 0.2% solution containing 1 mg free base per cc (1:1000) in ampules containing 4 cc

C Method of Administration Add 4-16 cc or occasionally more Levophed[®] to 1000 cc of 5% dextrose for I.V. use through Murphy drip bulb. Determine response to initial dose of 0.25-0.5 cc of diluted solution per 10 Kg body weight and then maintain flow at rate to maintain blood pressure (usually to 0.5-1 mm Hg). Adrenalin is a very potent agent and great care must be employed in its use. Do not allow solution to infiltrate or slough may result.

MALE SEX HORMONE

A number of natural hormones have been isolated from the testis and of these the most important androgen has been testosterone. It is believed that the effect of testosterone on the male sex hormone Testosterone is responsible for the development of secondary sex characteristics in the male (i.e. facial hair development, enlargement of prostate gland, seminal vesicles). Administration of testosterone to the female causes development of male secondary characteristics. In the female the androgenic effect is counterbalanced by the simultaneous administration of female sex hormone.

Oral administration of testosterone is androgenic effect is isopropyl and (the so-called) testosterone. Testosterone has mild androgenic effect and water retarding effect.

Oral administration of testosterone is androgenic effect is isopropyl and (the so-called) testosterone. Testosterone has mild androgenic effect and water retarding effect.

Testosterone

Testosterone is not active when swallowed. At present the only way to administer the agent effectively is to place it under the skin by injection or a pellet. A test on preparation which does not occur naturally in the testis (MT) is effective when swallowed. Methyltestosterone, a human androgen, is a marked androgen and has apparently produced jaundice after prolonged administration with a high rate of metabolism and androgenic effect similar to that of testosterone.

requirements by injection exceed 3 mg ($\frac{1}{20}$ gr) one additional pellet should be implanted [e.g. for a requirement of 5 mg ($\frac{1}{12}$ gr) per day by injection implant 6 pellets]
Duration of action 6-8 months

4 Deoxycorticosterone trimethylacetate 25-75 mg i.m. once a month

B Adrenal Corticosteroid (F) (act. N.N.D.) May be administered i.v. subcut. or i.v. Used in treatment of Addisonian crisis

Dosage 20-100 cc (5-20 dr) or more daily as indicated

C Lipo Adrenal Cortex Steril Solution (C.A.) Administered i.m. only

Dosage 5 cc ($1\frac{1}{4}$ dr) i.m. daily during crisis in addition to aqueous adrenal cortical extract 1-2 cc (16-32 dr) daily for maintenance

D Cortisone (Compound E) See page 424

E Hydrocortisone (Compound F) See page 424

F Fludrocortisone See page 425

G Preductal and Pred Solone See page 425

ADRENAL MEDULLA

Until recently it has been thought that the adrenal medulla secretes a single hormone epinephrine. However it has been shown that the tracts of adrenal medulla of cattle (*U.S.P. Reference Standard*) contain two closely related hormones i.e. epinephrine (about 80%) and nor epinephrine (about 20%). The two have different actions as outlined below

Substance	Blood Vessels	Cardiac Output	Blood Pressure	Blood Sugar (Glycogenolysis)
Epinephrine	Vasodilation (over all)	Increased	Elevated	Elevated
Nor epinephrine (levarterenol)	Vasoconstriction (overall)	No effect	Elevated	Elevated 1/3 that of epinephrine

Vasodilator of coronary arteries

Since epinephrine may be synthesized or derived from natural sources (usually the latter) and hence contains natural with nor epinephrine the reason for some of the apparent paradoxical physiological effects of the peptide preparation becomes clearer.

In addition to the above epinephrine causes immediate elevation of blood sugar by inducing glycogenolysis in liver and muscle.

Epinephrine

A Clinical Use Epinephrine is used in a great many clinical conditions including the following

- 1 Allergic conditions Bronchial asthma urticaria angio neurotic edema and others
- 2 Control of superficial bleeding especially from mucous membranes
- 3 Used with local anesthetics to slow down absorption

C Ch l f P p ratl In view of th g cal n mbe of p ep-
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FEMALE SEX HORMONES

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E t g ns

A Effect f E t og n in the Human The p im ry ctions of
est og n ar

- 1 P life ation f dom tr um
- 2 Changes in v gin l ell (co nfil ation and low ring of v gi
n l pH below 4 0)
- 3 D ct l prolife ation f b east
- 4 Stimulation of osteobl ti activity
- 5 Slight p ot in anaboli ffect
- 6 Mod ate od um and wate ret ining effe t

B Clini al U e E t ogen e us f l in both for their
f f ct on st blast n th tr tm t of osteoporosis

- 1 Fem le E trog ns us d a repla m nt the py in c acs
of ian fail e (g m opaus)
- 2 M le U d adjun t in th apy of ar troma of p ostate

C P ep rat ns Availabl The e a e many subetan es that hav
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and al o havu g th p utl and to l l vels that ar qu t im
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The is little on ed at p es t t admin te th e tro
ge e by any b t the o al out ab orption ms to be v ry
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be d cr as d by p nte al dmini t ation Th e is l k wi
no vid e th t th natu ally oc ur ing trogen a e any
mo ffic iou than the synthet on s Although st g n
pp ntly pl ya ol in mammary tum s of anim l th is
n vid n that th y ar ar in g nic in h nans

and testosterone propionate. Testosterone and testosterone propionate when injected (or swallowed) are partially (about 30-50%) excreted as 17-ketosteroids in the urine. Methyltestosterone is not excreted as 17-ketosteroid. In fact its administration will result in diminished urinary 17-ketosteroids due to diminished endogenous testosterone production.

A Clinical Uses In either sex testosterone may be indicated in any debilitating disease for its protein anabolic function. In addition there are certain uses specific to the different sexes.

- 1 Male As replacement therapy in failure of endogenous testosterone secretion (e.g. hypogonadism, male climacteric). Its use in impotence, angina pectoris, homosexual activity and benign prostatic hypertrophy is without benefit.

2 Female

- a Gynecological conditions: Functional uterine bleeding, endometriosis, dysmenorrhea, and premenstrual tension.
- b Disease of the breast: Advanced carcinoma, chronic cystic mastitis, suppression of lactation.

B Preparations Available

1 Testosterone USP (Free)

- a Pellet USP 75 mg implanted subcut. Dosage 4 pellets every 3-4 months.

- b Microencapsulation in aqueous solution for I.M. use. The dosage has not yet been determined but appears similar to testosterone propionate in oil.

Oil solution 5 mg/Gm for local testosterone effect. Average dose 0.5-1 Gm locally rubbed in once or 5 min b.i.d.

- 2 Testosterone Propionate USP B.P. I oil for I.M. injection 5, 10, 25 and 50 mg/. Dose varies from 10-100 mg daily depending on clinical indication.

- 3 Testosterone Cypionate USP B.P. (Depot testosterone) I oil for I.M. injection 50 and 100 mg/cc. This preparation has a duration of action 2 to 5 times or more than that of testosterone propionate. Dosage 100-200 mg weekly to 500 mg monthly as a single dose.

- 4 Testosterone Enanthate USP B.P. (Delatekyl®) in oil 100 mg/cc. The duration of action of this preparation is comparable to that of Depot testosterone. The average dose is 200-400 mg every 3-4 weeks.

- 5 Methyltestosterone USP B.P. Do not use methyltestosterone in treatment of thyrotoxicosis, acromegaly and glycosuria or liver disease.

Dosage 5-20 mg daily.

- a Tablets 5, 10, 25 mg for oral.

- b Tablet 5-10 mg for sublingual or buccal administration. There is no advantage of buccal over oral use.

Oil solution 5 mg/Gm.

- 6 Fluoxymesterone USP B.P. (Halotestin®) This drug is a fluorinated derivative of methyltestosterone. It is about 2½ times as potent as the parent drug. Its toxicity is less than that of methyltestosterone is not yet known. It is applied in 2 and 5 mg tablets. Dosage 1-2-10 mg daily orally.

- 7 Nandrolone (Nilex®) is a anabolic hormone with allegedly less corresponding androgenic effect than other testosterone preparations. It is applied in 10 mg tablets. Average dose 1-30-50 mg daily orally.

C Preparation Available See table below

Preparation	Method of Administration	Dose
Pegatron USP BP	IM	5-10 mg daily
Inject (Pegatron USP BP)	Pellet form plantion Oral tablets*	100-200 mg (for habitual or thrashed abuse)
Hydrocortisone Cortisone NND (Dilution)	IM 125 mg/c	125-150 mg every 2 weeks
Ethidione USP BP	Oral tablet	60-100 mg daily
Notethidione (No dilution)	Orally 5 mg tablet	20-50 mg daily (1 or two times a day as indicated)
The islet of Langerhans secretes corticotropin (ACTH) and the adrenal cortex secretes cortisones.		

CLINICAL USE OF CORTICOTROPIN (ACTH) AND THE CORTISONES

In the past few years both pituitary and corticotropin (ACTH) acting by adrenal stimulation and the cortisones have been shown to have profound modifying effects on many diseases. These effects are the result of the effects of these compounds on the basis of the known metabolic and immunological effects of these compounds (see page 411). These agents do not appear to cure. Their clinical appearance to be modification of cellular activity permissibility so that toxins no longer can affect the cell. When the drug is discontinued the disease may rapidly recur. No other hormones or combinations of agents that have a similar effect have the effect of these substances. The diseases in which ACTH and the cortisones have been tried in sufficient instances to warrant their evaluation are outlined on page 424. In general it can be said that these agents are interchangeable but occasionally a patient found to be responsive to one of these hormones and not to the other.

Dosage

Dosage varies considerably in each individual. In general it is advisable to employ the lowest dose that gives the desired effect. It is generally advisable to begin with larger doses and then decrease as rapidly as possible.

- Corticotropin USP (ACTH) May preparations (see pages 424 and 425)
- Cortisone Acetate USP Available under various names (see pages 424 and 425)
- Hydrocortisone USP Now available commercially Hydrocortisone is probably the principal steroid secreted by the adrenal cortex. It is about 40% of the half-time as potent as cortisone (see pages 424 and 425)
- Dehydrocortisone NND Dehydrocortisone NND This is a derivative of cortisone. It is secreted by the adrenal cortex. It is about 40% of the half-time as potent as cortisone (see pages 424 and 425)

A list of a few estrogens with the most useful dosage in terms of approximately equal physiologic effects is given in the following table

Drug	Dosage and Administration	Comment
Diethylstilbestrol U S P Stillboestrol B P Tablets 0.1 0.5 1.0 mg (1500 1720 180 gr.)	0.1 0.5 mg daily by mouth	Synthetic nonsteroidal estrogen. Cheapest. Excellent preparation.
Hexestrol N F Dienestrol U S P Bestrol N N D	0.2 0.5 mg daily by mouth	No advantage over diethylstilbestrol. Generally more expensive.
Ethyl Estradiol U S P Tablets 0.02 0.05 mg	0.02 0.05 mg daily by mouth	Synthetic estrogen. Excellent preparation.
Estrone U S P (Theelin)	1 mg 2-3 times weekly or 1000 IU/daily I M	Little data; prefer conjugated estradiols below.
Estradiol Valerate N N D (Delestrogen®) (in sesame oil)	10-20 mg I M every 2-3 weeks	Long acting estrogens.
Estradiol Benzoate Injection U S P B P (solutions in oil)	0.5-1 mg every other day I M	Estrogens for injection are rarely needed.
Estradiol Dipropionate U S P	2-5 mg 1-2 times weekly	Slightly longer effect than estradiol benzoate.
Estrogen Substances Conjugated N N D (Amnestogen®) Premarin® Coneston®)	1-25 2-5 mg daily I M or by mouth	Natural estrogen. Rarely indicated.

Progestones

The progestones are of rather limited use in medicine.

A. Effect

1. Leads to the secretory phase of endometrium. Progestone in the absence of estrogens does not have any significant effect on the uterine muscle; the uterus must be stimulated (proliferated) by estrogen before progestone can act.
2. Acin proliferation of breasts.

B. Clinical Use

The progestone is of rather limited value in clinical medicine. Their main uses are:

1. Production of a normal menstruation. Progestone may be used with estrogens to maintain a normal cyclical menstruation in women who otherwise cannot menstruate.
2. Medical and C. Another important progestone is to produce the so-called medical dilatation and urethral dilatation. It is a usually a test of adequacy of endogenous estrogen production. It is performed as follows: Give 10 mg progesterone I M daily for 5 days or 125 mg Hydrocortisone acetate N N D (Dilalut®) I M on day 6. If menstrual bleeding occurs within 10-14 days of the first dose, it indicates that adequate amounts of endogenous estrogens are being produced.
3. In some cases of habitual abortion.

C Preparations Available Set out below

Preparation	Method of Administration	Dosage
Progesterone USP BP	IM	5-10 mg daily
Injectable of Progesterone USP BP	Parenteral plantations Olethylate	100-200 mg (for bilateral or threatened abortion)
Hydroprogesterone Capitate NND (Delalite)	IM 125 mg /c	125-150 mg every 2 weeks
Ethinone USP BP	Olethylate	60-100 mg daily
Nethalene (Nalidine)	Olethylate 5 mg t b l t	20-50 mg daily (two to four times as per treatment schedule)
The following differences in effects are shown		
progesterone ethinone		

CLINICAL USE OF CORTICOTROPIN (ACTH) AND THE CORTISONES

In the past few years both pituitary adrenocortical (ACTH) and gonadotropin stimulation and the corticosteroids (cortisones) have been shown to have profound modifying effect on many diseases. The effect is an not to be explained at present in the basis of the known metabolic and immunologic activities of these compounds (see page 411). These agents do not appear to cure. Their action appears to be a modification of cellular activity permeabilitys that to maintain homeostasis. When the drug is discontinued the disease as may rapidly recur. No other hormones or combinations of drugs is that available immediately have the effect of these substances. The diseases in which ACTH and the cortisones have been tried in sufficient instances to warrant the evaluation are outlined on page 428. In general it can be said that these agents reintroduce ability but once patient is found to be responsive to one of these hormones and not to the other.

Dosages

Dosage varies on the basis of individualizing each patient. It is generally advisable to begin with larger doses and then decrease as rapidly as possible.

- Corticotropin USP (ACTH) Many preparations available (see pages 424-425)
- Cortisone Acetate USP Available under various trade names (see page 424-425)
- Hydrocortisone USP New available commercially Hydrocortisone is the principal steroid used by the medical profession about one half time as as potent as cortisone (see pages 424 and 425)
- Dehydrocortisone NND Dehydrocortisone NND These are derivatives of cortisone and hydrocortisone respectively have little effect due to the fact that they appear to be poor for the patient compounds because they inhibit

CORTICOTROPIN (ACTH) AND CORTISONES

Corticotropin U.S.P. (ACTH) Lyophilized Powder	Mode of Administration	Daily Dose	Remarks
	If V Administer in any I V fluid by slow drip For maximum effect give I V during entire 24 hour period. May use for 8-12 hours a day.	5-40 U.S.P. units	Maximum therapeutic effect No resistance develops if it does occur after I.M. injections I.V. administration is effective
Aqueous Solution	I.M. Administer in saline every 6 hours Administer as above either I.V. or I.M. Prepared by diluting with distilled water	40-200 U.S.P. units As above	Resistance often develops with continued I.M. use of this preparation Resistance may occur depending on method of preparation
Corticotropin Depot Repository Corticotropin Injection U.S.P. (Corticotropin Gel)	Give I.M. about every 12 hours for maximum effect. May use once a day in some patients	10-200 U.S.P. units	No development of resistance
Corticotropin Acetate U.S.P. Tablets 5 and 25 mg	Oral administration. Give in divided doses every 6 hours or q.i.d.	25-200 mg or more	Rarely used because of sodium retention. Rapid withdrawal may lead to hypoadrenal crisis. Slowly absorbed and slowly excreted.
Spermaceti (aqueous) 25 mg/cc	I.M. in doses every 12-24 hours	25-100 mg or more	For local use only
Eye Drops and Ophthalmic Solution 0.5% 10-25 mg	0.5 or 2-3% solution instilled every 2-6 hours	Variable	
Hydrocortisone U.S.P. Tablets 5, 10 and 20 mg	Oral administration. Give in divided doses every 6 hours or q.i.d.	20-200 mg (1-40 mg)	Similar to cortisone clinically and metabolically. About 1/2 clinical potency on a weight basis. It is in Addison's disease.
Eye Drops 0.5-2.5% (in ophthalmic solution) Oral Cortisone 0.5-2.5%	As indicated in the eye drops (see above). May combine with corticotropin or cortisone.	Apply locally as directed	For all glaucoma and inflammation. Many patients combine with cortisone.

Hydrocortisone 100 mg in 20% solution in 100 ml of 50% alcohol	1 V Add to 300 cc or more of saline 0.9%	100-200 mg I V	Emulsion of cortisone in oil. Must dissolve at least 300 cc of oil.
Hydrocortisone 100 mg in 100 ml of 50% alcohol	Dissolve in 10 cc water. Administer I V	100-200 mg I V	Water soluble form. For use in emergency. May be dissolved in any quantity I V if needed.
Hydrocortisone 100 mg in 100 ml of 50% alcohol	Inject intramuscularly. Do not dilute with anything. Dose depends on joint size. Do not inject into joint.	10-37.5 mg into joints	Of value in inflammatory conditions of the joints where inflammation is relatively localized.
Hydrocortisone 100 mg in 100 ml of 50% alcohol	For intravenous use only.	10-37.5 mg into joints	As hydrocortisone acetate. For systemic effect.
Prednisone 5 mg in 100 ml of 50% alcohol	Oral administration. Use as hydrocortisone.	5-50 mg or more (avg 10-20 mg)	Adrenal cortex equivalent. Does not have significant sodium retaining effect. About 4 times as potent as cortisone in inflammatory effect. Parent drug. Drug of choice.
Prednisone 5 mg in 100 ml of 50% alcohol	As hydrocortisone acetate. For intravenous use only.	10-37.5 mg into joints	Very effective. For intramuscular use.
Prednisone 5 mg in 100 ml of 50% alcohol	For intravenous use only.		As for hydrocortisone acetate (see above).
Prednisone 5 mg in 100 ml of 50% alcohol	For intravenous use only.		Very effective. For intramuscular use.
Prednisone 5 mg in 100 ml of 50% alcohol	For intravenous use only.		Very potent. Local use in myeloma. Systemic absorption with prednisone.
Prednisone 5 mg in 100 ml of 50% alcohol	For intravenous use only.		1 1/2 times as potent as prednisolone. No therapeutic advantages.
Methylprednisolone (Medrol)	Oral tablets	4-40 mg	No therapeutic advantages.
Do not give with specific diseases			

little if any sodium and water retention yet retain all other metabolic and therapeutic effects. Because of their lack of salt and water retention they must be used with even greater caution for the other hazardous effects of cortisone therapy are still present and a common early sign of overdosage edema may be absent.

- E Fludocortisone Acetate, N N D Usually used only topically (see page 425) or as supplement for maintenance in Addison's disease.

Duration of Therapy

It appears at present that prolonged administration will be necessary and can be employed safely in many cases. Where knowledge is available regarding recommendations for treatment these are indicated in the text.

Dangers

In severe agents are potentially very dangerous. However with proper caution most of these dangers can be overcome. The principal dangers are that these drugs may induce:

1. Hyperglycemia and glucosuria (diabetogenic effect). This is of major significance in the early or potential diabetic.
2. Marked retention of sodium and water with subsequent edema, increased blood volume and hypertension.
3. Negative nitrogen balance with loss of body fluid including bone protein with resultant osteoporosis.
4. Potassium loss with development of a hypokalemic alkalosis.
5. Hirsutism and acne (especially undesirable in females).
6. Cushing's syndrome (may develop with long continued administration).
7. Activation or production of peptic ulcer.
8. Lowering of resistance to infectious agents.

Control of Therapy Employed to Correct Minor Manifestations

- A Always reduce the dosage as soon as consistent with the clinical response.
- B During the first 2 weeks of therapy the following should be carefully observed:
1. Blood pressure
 2. Weight
 3. Initial complete blood count reported as indicated.
 4. Initial sedimentation rate reported as indicated.
 5. Urinary sugar, fasting blood sugar if doing substances are found in the urine.
 6. Serum potassium and CO_2 should be checked occasionally if large doses of hormone are to be given or more than several days.
 7. Daily eosinophil counts or measurement of urinary steroid excretion if a question of lack of adrenal response to corticotropin arises.
- C All patients should be on high protein diet (100+ Gm protein).
- D If edema develops place patient on low sodium diet (200-400 mg sodium daily). Mercaptopurine may be employed when strict sodium restriction is impossible.
- E Potassium chlorate 3-15 Gm daily in divided doses should be

- administered for 1 g daily or high dosage is employed
- F In cases of long continued administration it is to be one preparation (see page 420) if doses of 10-25 mg daily may be used to counteract the negative protein and potassium balance
- G Don't stop either drug abruptly. There may be a severe rebound of the disease process. Always remember that cortisone (hydrocortisone) causes atrophy of the adrenal cortex probably through endogenous ACTH inhibition. A sudden withdrawal may lead to symptoms of Addison's disease.

C. Indications and Special Precautions

- A Stress-Producing Agent Corticoid Patients receiving corticosteroids especially those in part of the body are carefully watched and managed. Because of the suppression of endogenous ACTH and subsequent adrenal cortical atrophy the patient is unable to respond normally to stressful situations (e.g., surgery, infection, etc.). Whenever such a situation occurs, a steroid should be given for corticosteroid hydrocortisone should be given and/or prednisolone corticoid and ACTH given. If corticosteroid or hydrocortisone is being administered it must be administered in large doses at least every 6 hours.
- B Hematology The age of the patient should be used with the treatment. In an individual with damaged myeloid marrow the intravenous or oral administration of a corticosteroid is recommended. Always begin with small doses and with the patient on low dosage.
- C Severe Rheumatoid These drugs probably contraindicated osteoporosis and should be used with caution in patients with major damage to the joints with dema and/or oliguria.
- D Prednisolone and Psychiatric These drugs cause a number of well-known side effects: most of these are weight gain but some individual (the predisposition to psychosis) may develop an acute psychosis, psychosis where the drug is discontinued. The patient should be watched for these and should be lowered and the patient should be fully observed and protected. Psychosis may be treated under the influence of the drug.
- E Effect of Thyroid When given for polypoid disease the drug may depress thyroid function. The majority inhibit the action of ACTH, the adrenal cortex depresses the action of corticosteroids in the cells of the body.
- G Effect of Prolonged Use Geoprolonged therapy of 65-90 mg (1.3 g) if the drug is to be given for more than 3-4 weeks.
- F Effect of Prolonged Use
1. At the point of use, a patient and at times of the drug, be aware of the danger of possible hemorrhage.
 2. Old patient or These get it difficult to activate. They should be given in the presence of the disease only as emergency with no only with optimum treatment with therapy.
- G Tuberculosis At times of the tuberculosis culture is so that it is to the effect of the drug.
- H Effect of Dose Because the drug is used to lower resistance and the top of the dose is of infection, they must be used with extreme caution when appropriate. Tuberculosis being given is a very useful and important.

RESPONSE TO CORTICOTROPIN (ACTH) AND CORTICOIDS IN VARIOUS DISEASES

Usually a Markedly Beneficial Effect

1 Rheumatic arthritis	10 Pemphigus
2 Acute rheumatic fever	11 Various allergic
3 Rheumatoid spondylitis	a Hay fever
4 Still's disease	b Angioneurotic edema
5 Psoriatic arthritis	c Drug sensitization
6 Acute gouty arthritis	d Serum sickness
7 Lupus erythematosus (early disseminated)	e Asthma
8 Inflammatory eye disease	12 Trichinosis
9 Exfoliative dermatitis	13 Nephrotic syndrome
(Only cortisone, hydrocortisone, and supplemental fludrocortisone are effective against Addison's disease and parathyroidism generally. Only prednisone and prednisolone are effective against the adrenocortical syndrome.)	
14 Ulcerative colitis (acute toxic)	

Results Encouraging but Often Only Transient

1 Regional enteritis	7 Pulmonary granulomas (except tuberculosis)
2 Periarthritis nodosa (early)	8 Toxic cytopnias
3 Scleroderma (early)	9 Chronic lymphatic leukemia (no longer responsive to other therapy)
4 Dermatomyositis	
5 Psoriasis	
6 Alcoholism (?)	

Transient Effect Only

1 Acute leukemia (lymphocytic or granulocytic)	4 Hodgkin's disease (rarely)
2 Multiple myeloma	5 Hepatitis especially hepatic coma
3 Lymphosarcoma	

No Beneficial Effects Established

1 Diabetes mellitus (increases insulin requirement)	8 Cognitive half-life (May be detrimental in some cases but case secondary to acute carditis may be beneficial though effect of hormone on inflammatory reaction)
2 Myasthenia gravis	9 Chronic myelogenous leukemia
3 Cash's syndrome	10 Poliomyelitis
4 Amyotrophic lateral sclerosis	11 Osteoporosis (may be detrimental)
5 Progressive muscular dystrophy	12 Acute monocytic leukemia
6 Progressive muscular atrophy	
7 Multiple sclerosis	

Chapter 17

NEOPLASTIC DISEASES

SOME SUGGESTIONS REGARDING MANAGEMENT OF PATIENTS WITH NEOPLASTIC DISEASES

Suggestions for Diagnosis

- A Early Symptomatology** When considering the diagnosis possible in patients with bizarre symptomatology remember that neoplastic diseases have an usually atypical phase in which they are less easily cognized than in more advanced clinical stages. *The importance of early diagnosis cannot be overemphasized.* Consider possibilities in the differential diagnosis of ill-unexplained illness especially in patients below 45 years of age. Do not convey this suspicion of neoplastic disease to the patient however unless it is necessary to discuss in detail the surveillance in completing diagnosis.
- B Physical Examination** In addition to detailed examination of the suspected areas physical examination must always include careful survey of the skin lymph nodes bimanual breast palpation and rectum.
- C Intramural Examination** Endoscopy gastroscopy sigmoidoscopy transrectal sonography of rectal lesions.
- D Special Laboratory and X-ray Studies** In addition to CBC urinalysis and total ammonia nitrogen the following are useful in the diagnosis of neoplastic disease.
- 1 Microscopic studies of smears and tissue sections
 - a Biopsy of accessible tumors and/or nodes
 - b Metastatic biopsy
 - c Cytologic examination by means of smears of all in body fluids (transudates and exudates) and body secretions (e.g., bronchial secretions)
 - 2 X-ray studies. Plain contrast studies of suspected areas if evident for primary or metastatic neoplastic disease.
 - 3 Chemical methods
 - a Blood hematology studies (e.g., blood lactate in pancreatic carcinoma)
 - b Catecholamine levels (e.g., hypothyroidism of gastrointestinal cancer)
 - c Blood and urine excretion tests (e.g., blood phosphatase in prostatic carcinoma)
 - d Blood hormone tests (e.g., blood protein bound iodine in thyroid disease)
 - e Hormone excretion tests (e.g., 17-ketotestosterone in adrenal tumor)

- E Surgical Exploration** (e.g. simple incision, thoracotomy or laparotomy) May be indicated as a final evaluation measure. In many cases the surgeon must be prepared to perform a radical surgical operation if macroscopic or frozen section examinations indicate malignant disease.
- F Reexamination** Upon completion of the clinical studies *emphatic reassurance of the patient regarding the negative findings is necessary*. If findings are equivocal the patient should be kept under close follow up observation with appropriate diagnostic measures.

Suggestions for Treatment

A Factors Influencing Choice of Treatment

1. Nature (inherent characteristics) of the given neoplasm: rate of growth, cytological characteristics, invasiveness, amenability (e.g. radiosensitivity or radiocurability), tendency to metastasize, and nature of metastasis.
2. Age of patient
3. Physical and emotional status of patient
4. Patient's ability and/or willingness to cooperate with the prescribed therapy
5. Availability of professional and technical facilities
6. Stage of the tumor at the time the patient is first seen
7. Location of the lesion: Proximity to vital or tubular structures
8. Secondary complications of the disease: Local pressure, symptom, hemorrhage, cystic effects of the primary lesion and the metastases
9. Functional, cosmetic and psychological effects of therapy
10. Patient's ability to tolerate radiation therapy (if tolerance of solar or other radiation)
11. Cost of therapy

B Treatment of Benign Lesions The physician's clinical impression of the benign character of lesions must always be verified by biopsy and microscopic examination.

1. Simple eradication of the tumor by surgical techniques (including urettag and cauterization) is usually the preferred method of treatment. Radiation technique may occasionally be employed.
2. General indications for removal of benign tumors:
 - a. Diagnostic purposes (possibility of malignancy)
 - b. Pressure on vital structures
 - c. Obstructive symptoms
 - d. Mechanical (static) deformities
 - e. Pain or other marked discomfort
 - f. Systemic effects (e.g. hormonal)
 - g. Hemorrhage (acute or chronic)
 - h. Cosmetic purposes
 - i. Psychological purposes (reassurance)
3. More extensive surgery: The surgeon must be prepared to perform radical surgery if macroscopic appearance or frozen section examination indicate malignant disease.

C Treatment of Malignant Lesions

1. Primary Lesion

- a. Complete eradication of the primary lesion by surgical

(including curettage and cauterization) or resection. Methods must be attempted whenever possible.

- b Radical surgery. Clinical evidence of regression in metastases may indicate need for radical surgical removal of the primary tumor and the involved nodes.
- c Surgical removal of the primary tumor may still be indicated when metastases are systemic but are growing very slowly (e.g., thyroid carcinoma).
- d Radiation therapy may be used to arrest or slow the progress of the disease if the tumor is radiosensitive.
- e Chemotherapy. See below.

2 Metastatic lesions

- a Surgical excision may be of value when lesions are solitary, slowly growing, painful, or when they produce other acute symptoms (obstruction, etc.).
- b Radiation therapy is indicated if lesions are radiosensitive and particularly if they are multiple or disseminated.
- c Chemotherapeutic methods may be employed using the specific agent which are known to affect certain types of primary and metastatic tumors. The agent is ordinarily withheld until there is definite need for symptomatic relief.

(1) Androgenic and estrogens. Definitive beneficial effects have been observed with the empiric use of the estrogens in certain neoplastic diseases but much of the work remains on an experimental basis. The duration of effect is unknown. Steroid therapy is never curative and should never replace radical surgery of operable carcinoma.

(a) Estrogens. The degree of the estrogenic response in individual cases depends on the patient's response and the toxicity of the drug (anorexia, nausea, diarrhea, dermatitis, and edema).

(1) Soft tissue metastases from bronchial carcinoma (i.e., lungs, breast, etc.). Tamoxifen improves mentorexia in a small percentage of elderly patients. In general, estrogens are reserved for patients 5 or more years past the menopause. Give Diethylstilbestrol (U.S.P.) 5-30 mg (usually 10-15 mg) or Ethinyl Estradiol (U.S.P.) 0.2-0.5 mg orally daily. Cyclical administration (i.e., 40 days on, 10 days off) is recommended.

(2) Pain in metastases (see page 309).

(b) Androgen. Methyltestosterone (U.S.P.) 5-10 mg sublingually daily or Testosterone Propionate (U.S.P.) 75-200 mg i.m. 3 times weekly per month may be indicated for:

- (1) Carcinoma of the cervix or ovary. Clinical complete relief of pain for variable periods but no objective improvement.
- (2) Breast carcinoma. Small metastatic lesions in 15% of cases show improvement but only occasional improvement is observed in soft tissue metastases.

- (2) Nitrogen mustards Although employed with benefit in certain cases of metastatic carcinoma these agents have proved most beneficial in certain diseases of the blood and lymphatic systems (see page 241)
- (3) Mustard like compounds (TEM TEPA) Similar to the above although less toxic (see page 238)
- (4) Antimetabolites (aminopterin 6 MP) See pages 233 239
- (5) Urethane® See page 239
- (6) Radioactive salts Effects are due to radiation rather than chemical action

S When none of the above procedures is possible

- a Narcotic drugs Liberal but judicious use especially in advanced and terminal malignant disease
- b Surgical measures

- (1) Relief of specific symptoms Surgical intervention (e.g. tracheotomy thoracentesis paracentesis lumbar puncture etc.) may be necessary to control progressive or emergency obstructive or other pressing symptoms
- (2) Nonspecific surgical methods (hormonal modification)
 - (a) Adrenalectomy Bilateral removal of the adrenal glands can sometimes produce a substantial regression of extensive and widespread male and female mammary cancer Although there is objective as well as subjective evidence of improvement the relief is most often of temporary nature This is still largely a research technique since it is a major operative procedure expensive and requires careful follow up steroid replacement therapy The use of this procedure in the treatment of other neoplasms is being investigated but no significant statistical results are available at present
 - (b) Ovariectomy Removal of the ovaries has been demonstrated for some time as a treatment for advanced breast cancer The result of the operation are usually transient and the relative efficacy of the treatment has been questioned
 - (c) Orchiectomy Castration may result in significant regression of primary and secondary tumors of the prostate and male breast In patients who fail to respond to orchiectomy subsequent adrenalectomy may prove to be effective Subjective and objective relief may be sufficient rather than a cure

D General Problems

1 Explanation to the patient

- a Factors of importance Opinion varies greatly as to whether or not it is advisable to inform patient that they have malignant neoplasia etc This matter must be individualized and must naturally vary with the temperament intelligence attitude and desires of the patient Under certain circumstances it may be necessary or advisable to inform the patient as to the true nature of his condition irrespective of the above factors
- (1) If the patient demands an explanation of his illness

- (2) If the patient's economic status requires forbidding to permit proper disposition of estate
- (3) If the patient refuses to carry through on a prescribed diagnostic and/or therapeutic regimen
- (4) If the neoplasm is growing relatively slowly, is non-invasive and is readily treatable
- (5) If the patient exhausts (or threatens to exhaust) his financial resources for a cure

b. Manner of explanation. If explanation is indicated use mild terms such as growth, lump, or even tumor, but in most cases it is advisable to avoid the term *cancer*. Be guarded in statements as to prognosis and lean toward the optimistic side. *Always offer some ray of hope.* When the clinical situation is not utterly hopeless, cheerful optimism and reassuring attitude may do much to allay the fear and apprehension of the patient and the family.

2. Explanation to the family. It is often advisable to inform a near relative (preferably the mother when this is feasible) of the nature of the illness and the prognosis. The qualifying statement should also be kept in mind in deciding who, how, and what to tell.

3. Provisions for chronic and terminal care.

a. Assessment of social significance. In view of the chronicity and the psychological and socioeconomic implications of the illness, the help of a medical social worker is desirable in proper fitting.

b. Hospital or nursing home. May be indicated.

c. Home. If patient and family decide on home care, it will be necessary to instruct some members of the family in the technique of administration of drugs (especially parenteral narcotic).

TREATMENT OF ADVANCED MAMMARY CANCER

	Premnopausal	Postmenopausal
<p>EXCISION OF GONADS (OVARIECTOMY)</p> <p>↓</p> <p>Eliminate ovarian function</p>	<p>Impaired fertility because of loss of ovarian function. This method is indicated in premenopausal women.</p>	<p>Estrogen therapy in postmenopausal women.</p>
<p>OVARIAN IRRADIATION</p> <p>↓</p> <p>Eliminate ovarian function</p>	<p>Best results are obtained when combined with systemic therapy and pulmonary irradiation (50%).</p>	<p>Less effective than in premenopausal women.</p>
<p>ANDROGENS</p> <p>↓</p> <p>Progestins (Mestranol)</p> <p>Ovarian progestins</p>	<p>Symptoms relieved 66% of patients and objective improvement in 20% of patients. Progestins prolong life.</p>	<p>Ineffective.</p>
<p>ESTROGENS</p> <p>↓</p> <p>Estrogens (Mestranol)</p> <p>Estrogens</p>	<p>Probably only 5% of patients benefit.</p>	<p>Subjective improvement in about 50% of patients and objective improvement in 25% of patients. Prolongation of life.</p>
<p>ADENAELECTOMY (BILATERAL)</p> <p>↓</p> <p>Eliminate production of androgens and testosterone (Pituitary and D.O.C.A. after operation)</p>	<p>Ineffective.</p>	<p>Subjective improvement in about 25% of patients and objective improvement in 10% of patients.</p>

METHODS USED IN TREATMENT OF NEOPLASTIC DISEASES

M. bud	M. h. m. (A)		S. G. K.		S. G. K.		S. G. K.		S. G. K.	
	R m	l f p im y	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m
ORIGINAL EXCISION										
	R m	l f p im y	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m	cont y m
RADIATION										
X y	D t	ti f p im ry	d y	d y	d y	d y	d y	d y	d y	d y
R dium	sum	i								
R dium ti S li	S i	s	d d	s	s	s	s	s	s	s
R diophosph ru	S i	s	d d	f m	f m	f m	f m	f m	f m	f m
R diocobi	S i	s	d d	f m	f m	f m	f m	f m	f m	f m
R d g id	S i	s	d d	f m	f m	f m	f m	f m	f m	f m
REMOVAL MODIFICATION										
A dr g na	P on	i an b li	ff t	iam pp	iam pp	iam pp	iam pp	iam pp	iam pp	iam pp
	lon (?)									
E t e	Emp	i	ff t							
C i	Emp	i	ff t							
	Elim	i	f col ti	f t	f t	f t	f t	f t	f t	f t
Ad	Elim	i	f col ti	f t	f t	f t	f t	f t	f t	f t
G i	Elim	i	f col ti	f t	f t	f t	f t	f t	f t	f t
G i	Elim	i	f col ti	f t	f t	f t	f t	f t	f t	f t
ANTINEUTROPHILS										
A i f li A id C mpoed	C	mpetition with (f l i d)	blocki g	f f	f f	f f	f f	f f	f f	f f
Amin pt i A M thopt i	ym h	i d	i s a id							
d Ad opt i	t m	g wh								
A ti sman A d C mpo und	C	mp i tion w h m i	d blocki g	y	y	y	y	y	y	y
2 6 Di m i P in	f ynth	i f m i n	i d							
8 M p i o p i (8 MP)	f t m	g ow h								
NITROGEN MUSTARD COMPOUNDS										
HN ₂ (M thyl bi (2 hi oethyl) m i HCl)	Dur	ti t i	ity t i m	ti	ti	ti	ti	ti	ti	ti
MUSTARD LIKE COMPOUNDS										
T i thyl m i m i n (TEM)	Dur	ti t i	ity t i m	ti	ti	ti	ti	ti	ti	ti
T i thyl m i m i n (TEPA)	Dur	ti t i	ity t i m	ti	ti	ti	ti	ti	ti	ti
U han	Dur	ti t i	ity t i m	ti	ti	ti	ti	ti	ti	ti
H i f (My)	D p	my i p i								

Chapter 18

VENEREAL DISEASES

SYPHILIS (Lues)

An acute or chronic disease caused by infection with *Treponema pallidum*. It may be either congenital or acquired. The acquired form of the disease is usually transmitted genitally but may be acquired by extragenital routes.

DIAGNOSTIC FEATURES

Primary Syphilis (code No 147)

- A History of contact with an infected individual 1-8 weeks (usually 3-4 weeks) prior to appearance of primary lesion
- B Primary lesions are pleomorphic; may be single or multiple and are usually located on the external genitalia although extragenital hauc are not rare (10-15%)
- C Three or more carefully performed dark field examinations (on successive days) are necessary before a final report of negative may be made
- D Both complement fixation (e.g. Kolmer) and precipitation (e.g. Kahn) tests should be performed. Quantitative blood tests (performed by a reliable laboratory) when they may demonstrate changing titer are preferred for both diagnostic and follow-up purposes
- E Regional lymph node on one or both sides are often rubbery discrete and nontender

Secondary Syphilis (code No 013-147)

- A Usually occurs 7-10 weeks after exposure to the disease and 2-3 weeks after appearance of the primary lesion
- B There is often evidence of systemic involvement with fever, generalized lymphadenitis, nonpruritic maculopapular dermatitis, nasopharyngitis, laryngitis, conjunctivitis, alopecia, arthritis, mucous patches and condylomata
- C Blood tests for syphilis are almost invariably strongly positive
- D Cutaneous and mucous membrane lesions may show *T. pallidum* on dark field examination
- E Spinal fluid usually shows transient involvement

Relapsing Syphilis

- A Usually occurs within 6 months to 2 years after onset of the disease

- B Often follows in adequate or improper therapy (e.g. penicillin for congenital gonorrhea)
- C Blood tests for syphilis usually revert to a positive reaction or if already positive to an increasing serologic titer (based upon quantitative blood tests)
- D Relapse may be of a several clinical type. The commonest of these is mucocutaneous. CNS, ocular and serological (the latter in the absence of clinical evidence)

Latent Syphilis (cod No. 400 147) (Early latent less than 4 years
late latent more than 4 years)

An intermediate or quiescent phase after secondary lesion has disappeared and while tertiary symptoms are not yet visible

- A Latent syphilis is a nonclinical evidence of infection than the positive blood test. It is therefore important to select the positive blood test, the most common auscultatory which are technical or clinical disorders acute fevers, yaws, infectious mononucleosis, malaria, leishmaniasis, smallpox, varicella, lymphogranuloma venereum, syphilis, lupus erythematosus, thrombocytopenia, and biological false positive reactions

Never make a diagnosis of latent syphilis solely on the basis of a single blood test. Rule out the possibility of the above factors. If the blood test is only very transiently and weakly positive the diagnosis of lues should be questioned. Conversely if the blood test is persistently positive for 3 or more months lues is the most likely diagnosis.

- B Spinal fluid must be completely negative
- C The treponema immobilization test (TPI) and the Treponema pallidum complement fixation test (TPCFT) although not available for routine use will distinguish biological false positive reactions from true syphilitic reactions and should be employed in all instances of doubtful diagnosis

Late (Tertiary) Syphilis (cod No. 414 147)

It is characterized by diffuse, mainly cutaneous, ganglionic, and may be localized or disseminated lesions (gummas) in any and all tissues

- A Mucocutaneous Gummatous lesions of the skin and mucous membranes

- B Osteo Diffuse or gummatous lesion of bones and joints with periostitis, arthritis, synovitis and osteomyelitis

- C Ocular Conjunctivitis, iritis, vitritis, choroiditis, keratitis and hemorrhagic

- D Visceral (excluding cardiac) Gummatous or diffuse involvement of pancreas, lung, spleen, kidneys and testes

- E Cardiovascular

1 Uncomplicated aortitis (cod No. 461 147)

2 Aortic aneurysm (cod No. 455 147)

3 Aortitis (cod No. 461 147 8)

- F Neurosyphilis

1 Asymptomatic neurosyphilis characterized by spinal fluid abnormalities (Goussard's page 438) but without evidence of symptoms or signs of neurological involvement

Spinal Fluid Findings in C N S Syphilis*

Group and Degree	W b c per cu mm.	Pandy's Test	Total Protein (mg per 100 cc)	Complement Fixation Test	Colloidal Gold Test
I Mild or minimal	8 or more	- or ±	5-50	- or ±	0000000000 to 2 21000000
II Intermediate or moderate	8-200 or more	± or +	40-100	± or + with 1 cc of fluid	0012210000 to 3332100000
III Severe or maximal (paretic)	8-200 or more	± or +	40-200	+ with 0.1-0.5 cc of fluid (strongly positive)	55543 1000

Spinal fluid must be non-bloody

- a May be classified according to C S F changes (see above) as mild moderate or severe (Groups I II or III)
 b If untreated may develop into clinical neurosyphilis
 May occur during any phase of life

2 Symptomatic

- a Acute syphilitic meningitis (code No 912 147 0)

- (1) Usually occurs within 2 years after infection
 (2) Clinical picture of low grade meningeal irritation
 (3) Spinal fluid Group I and II changes
 (4) Often follow inadequate treatment

- b Chronic syphilitic meningitis (code No 912 147)

- (1) Usually occurs 2-20 years after infection
 (2) Clinical picture varies considerably according to portion of C N S involved
 (3) Spinal fluid Group I and II changes (see above)

- c Diffuse meningovascular (code No 910 147)

- (1) Involve both meninges and blood vessels Vascular thromboses are frequent
 (2) Clinical picture varies with C N S location of thromboses and includes both neurological and psychiatric manifestations Repeated thromboses may occur
 (3) Prognosis is usually good with treatment
 (4) Spinal fluid Group I and II changes occur (see above)

- d Tabes dorsalis (code No 906 147)

- (1) Usually occurs 2-20 years after infection
 (2) Involves dorsal spinal cord and columns midbrain base of pons and autonomic nervous system
 (3) Clinical picture includes ataxia pains of varying character and location visual disturbances phosphenes and sexual disturbance pupillary changes optic atrophy hyporeflexia diminution of vibration and position sense and other sensory disturbances
 (4) Spinal fluid Group I and II changes occur in the earlier stages of the disease changing little to Group III abnormalities although the C S F may be completely normal (see above)

(5) Blood test for syphilis are positive in 50-75% of cases of tabes

e Taboparisis (code No 906 147 9) Combines clinical features of tabes and paresis. Spinal fluid is variable

f General paresis (dementia paralytica) (code No 00 147)

(1) Usually occurs 2-20 years after infection

(2) A syphilitic meningoencephalitis

(3) Clinical picture variable but may include mental deterioration, personality and behavior disturbances, convulsions, paralysis, weakness, tremors, pupillary changes, speech defect, hyperreflexia and other evidence of motor neurone involvement

(4) Spinal fluid Group III changes occur (see page 438)

(5) Blood tests for syphilis are positive in 50-100% of cases of paresis

g Optic atrophy (code No 962 147 9) Disc pallor, visual field disturbances and altered visual acuity which in untreated cases progresses to blindness

Congenital Syphilis (code No 010 1471)

The clinical manifestations of the congenital disease are quite similar to those of the acquired form except for the atheromatous lesions, the absence of primary or initial lesions

A Evidence of fetal history of disease

B Skin and mucous membrane lesions at birth or in early infancy

C Characteristic stigmata of congenital disease such as interstitial keratitis, Hutchinsonian triad, eighth nerve deafness

(Hutchinsonian triad) saddle nose, hager's sab shin and the bone changes mentioned to

D Blood test for syphilis usually strongly positive at birth but gradually become negative over a period of years

E Any of the tertiary sequelae of the adult disease (CNS, vascular or digestive) may occur

TREATMENT OF SYPHILIS

General Measures

A Public Health Measures

1. In cooperation and sexual promiscuity people take with respect to sexual health should be somewhat isolated, quarantine until and then infectious by primary antilithic therapy
2. Report the infection to appropriate public health agency

B General Management

1. Completely rest, physical status of patient prior to institution of prophylaxis

A. Full physical examination, chest x-ray and fluoroscopic roentgenography. Initially a blood count are divided by the institution of antilithic therapy. Spinal fluid should be examined in all patients with latent or late disease prior to therapy

2. Mental disturbance, general health by adequate diet, rest, exercise and attention

C Local Measures (muco-cutaneous)

1. Local treatment is unnecessary

- 2 No local antiseptics or other chemicals should be applied to a suspected luetic lesion until repeated dark field examinations have been made. If after the diagnosis has been established the luetic lesion should become secondarily infected the lesion may be treated as for any pyogenic ulceration (this in addition to systemic antiluetic treatment see page 84)

Prophylaxis

- A Sex Education Instruction along the lines of sexual education is to be desired. Avoidance of illicit sexual contact is the surest of all prophylactic methods.
- B Mechanical The standard rubber condom is effective but protects covered parts only. The exposed parts should be washed with soap and water as soon after contact as possible. This applies to both sexes.
- C Antibiotic If there is known exposure to infectious syphilis abortive penicillin therapy may be used. Give 1,200,000 units of repository penicillin I.M. in one dose.

Recommended Treatment Schedules for Various Forms of Syphilis

- A Primary and Secondary Adult Syphilis Repository penicillin 600,000 units I.M. daily for 10 days (6,000,000 units).
- B Infectious Relapse Treat again as for primary or secondary syphilis. Diagnosis is made if careful follow-up examination reveals sustained or rising titer on serial quantitative biologic or monthly blood tests or if there is actual clinical evidence of relapse (mucous membranous lesions of mouth and ano-genital regions and skin lesions especially on palms and soles).
- C Latent Syphilis There are no clinical manifestations in latent phase and the C.S.F. is negative. The only positive criterion for this stage of the disease is the positive blood test. Only a small percentage of these blood titers however will be appreciably altered by treatment with penicillin. The treatment of this stage of the disease is intended to prevent the late sequelae.
- D Asymptomatic Neurosyphilis Benign Late Syphilis. Visceral and Cardiovascular Syphilis. Optimal dosage schedules of penicillin have not been completely established in these stages of the disease. Hazards of penicillin therapy (Herxheimer's reaction or therapeutic paradox) are minimal but the ultimate outcome of treatment is difficult to determine. The following dosage schedule is recommended. Repository penicillin 600,000 units I.M. daily for a total of 12,000,000 units.
- E Symptomatic Neurosyphilis
 - 1 Prognostic Neurosyphilis by establishing early diagnosis and by providing adequate treatment and follow-up of early syphilis. Examination of all syphilitic patients for evidence of nervous system involvement must be a regular part of the follow-up examination.
 - 2 The proper treatment clinical and laboratory evaluation should include detailed neurological, ocular and psychiatric examination and a cerebrospinal fluid examination. The high rate of coexistence of cardiovascular and C.N.S. lesions should be considered.

3 Treatment method

- a Penicillin treatment Penicillin is the treatment of choice in the syphilis. *Repen* 12 000 000 units I.M. daily to a total of 12 000 000 units may be given
- b Other methods The various schedules of metallotherapeutic drugs will be omitted here because they have become obsolete

- 4 Follow-up examination All patients must have a spinal fluid examination throughout the following completion of antileptic therapy. The general adequacy of response to treatment is at times difficult to evaluate (especially during a short period of observation) but it may be gauged by clinical improvement and effective adjuvant treatment of C.S.F. changes. A second course of penicillin therapy may be given if necessary

F Prenatal Syphilis

- 1 Make it a to expectant patient who are luetic the urgent necessity for antileptic therapy. Then make certain that appropriate treatment is carried out
- 2 Immediate treatment is important. Dosage schedules as advised for primary and secondary syphilis are satisfactory. When the spirochete is instituted late in pregnancy (i.e. after the 7th month) in women with untreated early syphilis the larger dose of penicillin is advised. Dosage schedules as outlined for symptomatic neurosyphilis and for cardiovascular syphilis are recommended. Remember that penicillin treatment may bring about cures in more than 90% of cases even when syphilis is discovered in the latter trimester of pregnancy
- 3 Follow-up must consist of monthly physical examination and quantitative blood serological test for syphilis (S.T.S.) until and for a month after delivery. If there is a significant fall in the S.T.S. titer, it should be pointed out to be expected that the result will necessarily be a satisfactory serological reactivity in month with late latent syphilis by any test method. If the mother has previously undergone treatment during the early latent syphilis and the original S.T.S. titer does not significantly decline within 3 months after treatment, treatment is advisable
- 4 The newborn infant should be examined for stigmata of flu and should be held to 2 to 3 weeks intervals for 4 months. If the maternal blood is positive, positive cord blood S.T.S. is a diagnostic aid. However, if the infant's blood is followed serially by quantitative blood S.T.S. at 2 week intervals for 4 months, a sustained or rising S.T.S. titer would indicate a diagnosis of congenital lues and a need for treatment

Special Consideration Regarding Penicillin Therapy

- A. *Herxheimer's reaction* is a familiar danger to encounter with marked fever and constipation and generalised ache and pain within 24 hours after onset of the spirochete
- B. Some clinicians feel that in late syphilis it is necessary to administer a course of bismuth and iodide prior to penicillin

therapy in order to diminish the hazard of Hxheimer's reaction or therapeutic paradox. These dangers if they exist at all are minimal.

- C Sensitivity to penicillin (see page 505) contraindicates further use. One of the other antibiotics may be given (see below).
- D Relapse following one or more courses of penicillin therapy requires consideration of other therapeutic agents.

Chlortetracycline (Aureomycin®) Treatment Methods

Oral Chlortetracycline Hydrochloride U.S.P. (Aureomycin®) has been reported to be effective in the treatment of syphilis but clinical experience with the drug is not extensive. Optimal dosage schedules, toxicity, failure rates, etc. remain to be determined. One Gm. every 4 hours day and night for a total of 70-80 Gm. has been suggested. Its use may be considered in those patients sensitive to penicillin. Tetracycline U.S.P. (Achromycin® Tetracycl®) may be used instead of chlortetracycline in a similar dosage.

DIFFERENTIAL DIAGNOSIS OF VENEREAL DISEASES

Disease Organism and How Demonstrated	Test	Lesions	
		Bubo	Genital
Syphilis <i>Treponema pallidum</i> (Dark field exam)	Complement fixation (e.g. Kolmer) and precipitation (e.g. Kahn) tests	Non-fluctuant	Painless ulcers
Chancroid <i>H. morph. ducreyi</i> (Gram stain)	Skin test with Ducrey antigen	Usually fluctuant	Painful ulcers
Lymphogranuloma venereum virus (Culture methods)	Complement fixation tests Frei test	Usually fluctuant	Painless venous ulcers
Granuloma inguinale Donovan bodies? (Wright stain)	Non	Usually none	Painless spreading ulcers
Gonorrhea (Gram stain) Nisseria gonorrhoea	Complement fixation (value?)	None	Urethritis

GONORRHEA

Gonorrhea is an acute or chronic infectious disease caused by the gram-negative diplococcus *N. s. s.* gonorrhoeae and practically always transmitted among adults by sexual intercourse. Acute gonorrhea in the adult male is characterized by acute urethritis with painful urination and purulent urethral discharge. Chronic gonorrhea may be manifested by chronic inflammation of the urethra, prostate, epididymis and seminal vesicles but rarely involves the upper urinary tract. Gonorrhea in the female begins in the urethra, vagina and vaginal glands and is characterized by painful urination and purulent discharge. Commonly the infection spreads to the uterus, tubes and other pelvic structures causing abdominal pain and is accompanied by constitutional symptoms. Systemic infection with septicemia and manifested by endocarditis or arthritis is less common. The disease is a stron-

affinity for the ocular mucous membranes and may cause a serious and blinding ophthalmia

Diagnosis

A History of genital discharge and dysuria occurring 4 to 10 days following sexual intercourse with an infected individual Symptom will vary with the anatomic structures involved Demonstration of the gram negative intracellular diplococcus in exudate from lesions by staining (Gram's or methylene blue) smear and by culture Bacterial culture may be positive in patients with purulent Copmplement fixation tests may be positive several weeks after the initial infection

B Microbiological Treatment Involvement

- 1 Urethritis (Code No 744 103)
 - a Smear and if necessary culture of material obtained from the urethral meatus will demonstrate the causative organism This is obtained by urethral tripping and never by any other method during the acute phase The discharge is unilateral in the chronic phase
 - b Two glass test tubes for first glass cloudy the second is clear Both glasses contain shreds Clinically symptom remits after a week and defecation and micturition may be normal in 1 day
- 2 Proctitis (Code No 764 103) Urethral discharge may or may not be increased Same smear and culture findings will be observed Penicillin is common and indicated by defecation Lower back pain may be present Constitutional symptom usually fever and chill may be present Dysuria is frequent and tenesmus may occur
- 3 Acute epididymitis (Code No 756 103) History of return urethral smear and culture findings as above There is also testicular pain swelling warmth and tenderness

C Femoral Gland Involvement

- 1 Acute urethritis (Code No 740 103) Smear and (preferably) culture of urethral and genital discharge should be performed There is redness and swelling of genital tubercle and external meatus
- 2 Chronic gonorrhea infections (after 4 to 6 weeks)
 - a Very fulminating prostatic cultures and titrated amount of intrabacillary infection from Skirrow's broth medium or endocervical glands (For treatment see U.S.P.H.S. VD Bulletin 97 1945)
 - b Pelvic inflammation (Code No 066 103) may be heralded by lower quadrant abdominal tenderness mild prostatic irritation and systemic manifestations

Treatment

Penicillin is protomyicidal in the acute phase (Aurum only in the acute phase) (Tetracycline[®]) and the sulfonamides are all effective against gonococci although penicillin is generally the drug of choice

A Acute Chronic Urethritis and Urethral Infection (male or female) (Code No 740 103) Avoid all local treatment such as irrigations manipulations and instillations

- 1 Penicillin therapy Several effective technics are available ALWAYS draw preliminary blood specimen for serological test for syphilis and examine patient clinically for evidence of syphilis since the danger of masking early syphilis by penicillin treatment is very real Give repository penicillin 600 000 units I M on 2 successive days
 - 2 Alternate therapy If coincident lues is suspected the above treatment should be altered as follows
 - a Penicillin Repository penicillin 600 000 units I M daily for 10 days
 - b If the patient is allergic to penicillin give one of the tetracyclins 1 0 Gm orally Stat then 0 5 Gm at 6 hour intervals for 4 6 doses
 - 3 Follow up Should consist of examination of the patient at weekly intervals for at least 3 weeks or preferably
 - a Weekly examination for evidence of urethral discharge chancre or rash
 - b Stained smear and if possible culture of any inflammatory exudate weekly Avoid prostatic massage urethral swabs or instrumentation as a means of obtaining material for examination in acute cases
 - c Blood test for syphilis and examination for clinical evidence of lues at the end of the third week and again at 3 6 12 and 24 months
 - 4 Retreatment of penicillin failures (suspect other etiology) If any of the weekly checks shows bacteriologic evidence of persistent gonorrheal infection repeat penicillin treatment as above Consider possibility of serologic complications If such can be reasonably excluded may treat with
 - a Increased dosages of penicillin
 - b Streptomycin sulfate U S P 0 3 0 5 Gm I M as a single dose
 - c Chlorotetracycline Hydrochloride U S P (Aureomycin®) 1 0 Gm orally Stat and the 0 5 Gm at 6 hour intervals for 4 6 doses
 - d Oxytetracycline U S P (Tetracycline®) 1 0 Gm orally Stat and 1 0 Gm repeated in 6 hours
 - 5 Persistent failures Often so called treatment failures are actually reinfections this becomes the problem if large has come to believe that modern treatment methods have now rendered the diseases virtually eradicated The danger of such a concept must be made apparent to the patient
- B Acute and Chronic Prostatitis (code N 764 103) Treat as above Hot sitz bath and alkalization of the urine may provide symptomatic relief
- C Acute Epididymitis (code No 756 103) Above treatment in addition to
- 1 Bed rest
 - 2 Cold compresses to scrotal region
 - 3 Analgesics for relief of pain
 - 4 Suppositories to be used during convalescent ambulatory phase
- D Pelvic Inflammatory Disease (Acute Gonococcal Salpingitis code No 787 103)
- 1 Absolute bed rest

- b Do douches or unnecessary manipulation during acute phase
- c Examine carefully for clinical evidence of lues. Draw blood for serological test
- d Repository penicillin 600 000 units I M daily for 3 10 days
 - (1) Convalescent period. If patient becomes afebrile and asymptomatic keep her at bed rest until WBC and sedimentation rate become normal (may take a month). Observe the patient during and following her next menstrual period for pain and pelvic change. If these are absent discharge her to home care on the convalescent program outlined below
 - (2) Retreatment. If symptoms fever leukocytosis increased sedimentation rate or positive vaginal smear persist or if they recur at the time of menses administer a second course of penicillin
- e Retreatment. If the patient fails to respond to 2 courses of penicillin therapy give one of the tetracyclines 1 0 Gm orally Stat then 0 5 Gm at 6 hour intervals for 4 6 doses
- f Convalescent program. After the patient is discharged from the hospital give the following instructions
 - (1) Sedentary life
 - (2) No sexual intercourse until signs and symptoms have completely cleared (usually takes about 6 8 weeks)
 - (3) Douche. Prolonged douches of warm tap water using 1 2 gallons and administering slowly and gently over a 15 20 minute period 2 or twice daily. The patient can perform this procedure most effectively while sitting in the bathtub
2. Subacute (or acute exacerbation of chronic) form
 - a Absolute bed rest until signs and symptoms have cleared
 - b Douches as above
 - Penicillin is much less effective in this phase of the disease but trial of therapy is warranted (see above)
3. Chronic (chronic gonococcal proctitis) (code No 787 103 0)
 - a Bed rest during exacerbations
 - b Penicillin usually ineffective but hold betide
 - Other antibiotics hold aloft
 - c A course of pelvic diathermy treatment may be of value
 - d Surgical procedures may be dictated. This decision should be made by a gynecologist. Result of surgery or of any therapy

GRANULOMA INGUINALE (code No 146 199)

A chronic granuloma disseminated by infection with *Dona* *g* *anul* *mat* *is* *a* *pl* *u* *m* *phic* *od* *it* *2* μ *in* *l* *g* *th*. It is manifested by painful sharply defined reddish ulcers and ulceratively bleeding granulomatous ulcers of the skin or mucous membranes of the genital region. If untreated the lesion gradually extends to involve the thigh and later the adjacent areas of the abdomen and thighs. It rarely involves deep structures. There is usually no lymphadenopathy. Lesions are

heal spontaneously although the process may remain stationary for years. Deep tissue scrapings or punch biopsy of clean peripheral granulation tissue should be stained with Wright's stain and examined for Donovan bodies (see Table on page 442).

Treatment

- 1 Chlorotetracycline Hydrochloride U.S.P. (Aureomycin®) Oxytetracycline U.S.P. (Terramycin®) and Chloramphenicol U.S.P. (Chloromycetin®) are all effective. 1 Gm daily for 12 weeks may be tried.
- 2 Streptomycin Sulfate U.S.P. is highly effective but may be contraindicated by the danger of damage to the vestibular apparatus. If tried it may be used in a dose of 1 Gm 1 M daily until the lesion is healed (10 or more days).

LYMPHOGRANULOMA VENEREUM (code No. 55-198) (Lymphogranuloma Inguinale or Lymphopathia Venereum)

Lymphogranuloma venereum is an acute or chronic venereal disease caused by a specific virus. It is characterized by minimal herpetiform genital lesions and may be complicated by regional lymph node involvement and at times by variable constitutional reactions.

The incubation period is one to three weeks. Initial lesions are painless to unnoticed herpetiform or ulcerative and may appear on any part of the external genital area. Inguinal buboes appear 1-4 weeks after infection, are often bilateral and may or may not be suppurative. The nodes may fuse, soften and break down, forming multiple sinistral tracts. Extensive arrhythmia may occur. Chronic anorectal disease manifested by rectal pain, sanguinopurulent discharge and rectal strictures is more frequently encountered in male than in female patients. Constitutional reactions frequently accompany the stage of bubo formation and are characterized by fever, chills and prostration, malaise and neurologic manifestation. Consider this disease as a possibility in undiagnosed cases.

The Frei skin test (chick embryo antigen) is of value. If suspicious lesions are of 3-4 weeks duration a negative result probably rules out lymphogranuloma venereum. A positive skin reaction may mean an active infection past (old) infection or related viral infection (false positive). The electrolysis of the albumin globulin ratio in the serum. Complement fixation test are valuable. Always rule out the possibility of primary syphilis (see page 436). False positive (usually weakly positive) blood tests for syphilis may occur.

Treatment

A. Specific Therapy

- 1 The tetracyclines and Chloramphenicol U.S.P. (Chloromycetin®) 0.25-1.0 Gm (3 3/4-15 g) orally q.d. for 5-14 days are the treatments of choice. Sulfadiazine U.S.P. or Sulfathiazole N.F. 1.0 Gm (15 gr) t.i.d. for 2-3 weeks or longer probably has no effect against the virus but is effective in preventing secondary complications.

B Local Measures

- 1 Bed rest (provides local comfort)
- 2 Warm fomentations to bubo perineal discomfort
- 3 Analgesic perineum
- 4 Aspirate fluctuant nodules under aseptic precautions (see below). Incision and drainage are to be avoided (to prevent lymphatic obstructions)
- 5 Proctoscopic examination for diagnosis and for late evaluation of changes
- 6 Extensive plastic surgical repair operations may be necessary in the chronic and rectal form of the disease. Rectal stricture should be treated by prolonged gentle dilation. Although in extreme cases this may be impossible and colon shunting procedures may be necessary.

CHANCROID (Soft Chancre)

(Of Penis code No 751 10x) (Of Vulva code No 774 10x)

A venereal disease caused by *H. morphilus ducreyi* and manifested by painful genital ulcer or ulcers often complicated by suppurating inguinal lymph nodes (buboes). Incubation period is from 3 to 5 days (range 2 to 7 days?) following venereal exposure. The inflammation begins as a small ovoid vesicopustule which ruptures to produce a shallow necrotic undermined ulcer. Single or multiple painful lesions may occur and phimosis may result. Regional lymph nodes become enlarged in a few days to weeks and are usually malleable, soft, fluctuant and tender. The nodes may rupture or may abscess spontaneously. Giemsa stained smears from the ulcer show *H. morphilus ducreyi* which may also be cultured from pus from the lesions of the bubo. Syphilis must be excluded by the diagnostic measures outlined under the diagnosis of syphilis (primary). The two diseases may coexist.

Treatment**A Specific Therapy**

- 1 Sulfadiazine U.S.P. or Sulfisoxazole U.S.P. (Gantrisin®) 1.0 Gm (15 gr) q.i.d. for 1 week. Observe for side effects with the following monographs (see page 501)
- 2 Chlorotetracycline Hydrochloride U.S.P. (Achromycin®) or Oxytetracycline U.S.P. (Tetracycline®) 0.5 Gm every 6 hours for 5-7 days

B Local Therapy

- 1 Careful irrigation with soap and water bid (after diagnosis has been made) will suffice. When lesions fail to heal promptly a saturated calomel ointment of 10-000 potassium permanganate solution may be necessary.
- 2 Fluctuant buboes may be aspirated with a large gauge (No 16) needle as indicated. Warm compresses or hot water bottle may be applied to the groin for relief and to hasten fluctuation and regression of bubo.

Chapter 19

INFECTIOUS DISEASES

DISEASES DUE TO VIRUSES

MEASLES (Rubeola) (code No 010 169)

An acute highly communicable virus infection characterized by inflammation of the respiratory tract conjunctivitis Koplik's spots and a blotchy rash.

The prognosis is generally good. Secondary infection by bacteria is common but responds readily to appropriate treatment. The fatality rate of post-measle encephalitis is 30 per cent and those surviving frequently have residual damage.

Diagnosis

Incubation period 10 days. A prodromal period of fever, cough and conjunctivitis precedes the rash by about 4 days. Koplik's spots usually appear 2 days before the rash. A blotchy rash appears on the face on the first day, spreads to the trunk on the second and to the extremities on the third and fourth days. Leukopenia present.

Measles is most contagious just before and during the prodrome. It remains contagious but less so for about a week after the appearance of the rash.

Treatment

A Specific Measure None available

B General Measures

- 1 Isolate for the week following onset of rash
- 2 Bed rest until afebrile
- 3 Aspirin as necessary for analgesia
- 4 Slight eye poultices for symptomatic relief of conjunctivitis
- 5 Vasoconstrictor nose drops
- 6 Sedative cough mixture if necessary (see p 110)

C Treatment of Complication

- 1 Secondary bacterial infection of the middle ear, throat, larynx or lungs are treated with appropriate sulfonamides or antibiotics (see 514)
- 2 Post-measle encephalitis (code No 930 169) may only be treated symptomatically
 - a Lumbar puncture for relief of headache, pressure
 - b Anticonvulsants as necessary (see 351)

P_{rophylaxis}

A ti e prophylaxis is n t pract cal but passive p otection or modification may be accomplished

- A C mple te tempo ary prote tion of xpos d s ceptibles usually f llow dm nst t o b e f re the sixth day of incubation o 20 cc of conv lescent serum 2 to 10 cc of immun serum globulin (gamma glob lin) o 3 to 10 cc of human immune gl bulin (pla ental immun) I M
- B Modif cation of the disease followed by p mane t immunity usually ults from the inj tion of half th above doses on th fifth t seve th days qual dos s o the eighth day or d ble doses o the ninth o tenth days following exposure

RUBELLA (German Measles) (code No 010 165)

R bell is an a te mmunicable dise s of viral origin charact ed by ra h and lymphad nopathy

Diagno is

Th in batio p riod i 2 to 3 weeks A short p odromal period of malais o a hi g in the post ior cervical nodes may p e e de the fi p pula erupts which appears usually fi st n the f e and quickly sp e d to the t unk and ext m ties S boc pital and p st ri cervical ad nitis is ually p esent Leukop nia is ge rally noted Pate ts a probably infectious d ring th p o d m and du ing the e ptio

T atm t.

- A Specific Mea s None
- B G I M s Asp in fo m lai If q i d
- C T atm t of C mpli at ons
- 1 Fetal bno m lity f q e tly found if the dis a oc urs d ing the f t o ly in the econd trim t of p g n y (S Prophylaxis)
 - 2 En ph iti (cod N 930 165) d th ombo ytopenic p r (cod No 516 165 9) e ve y ra Tre t sympt m tically
 - 3 S o d ry t pto oc al inf t on may occur a d ho ld be t t d with p illin (p 481)

P_{rophylaxi}

P gna tw m n who h ve b xp d to b lla may b giv n S 20 f immune s rum globuli (gamma globulin) I M in an ff rt to p ev nt o m dify the di

VARICELLA (Chickenpox) (code No 010 161)

Varic lla is an a ut communi able di a us d by a virus kin to that of h rp t It i ha act ri d by th e uption of ops of skin l ion

D gno i

Th in batio p riod i to 3 weeks (usually 17 day)

Prodromal symptoms are usually slight and last only one day. Lesions erupt in crops and progress through the maculopapular, vesicular, and pustular stages to crusts in about 3 days. The eruption is usually centripetal in distribution. The patient is infectious for one day before the onset and for 6 days thereafter. The late crusts may also occasionally be infectious.

Treatment

A Specific Measures None available

B General Measures

1. Isolate until primary crusts have disappeared
2. Bed rest until afebrile
3. Cleanliness of skin by frequent tub baths or showers when afebrile
4. Calamine lotion locally and antihistaminics orally may relieve the pruritus

C Treatment of Complications

1. Secondary bacterial infection of the lesions may be treated with bacitracin, tyrothricin, or penicillin ointment locally. If extensive, penicillin I.M. may be given.
2. Post-varicella encephalitis may be treated only symptomatically.

Prophylaxis

Temporary passive protection is regularly followed by I.M. administration of 20 cc. of convalescent serum, but this is rarely warranted.

SMALLPOX (Variola) (code No. 010 176)

Smallpox is a serious communicable febrile disease characterized by rapid onset of constitutional symptoms followed by an eruption most marked on the face and extremities and often involving the mucous membranes.

The prognosis is extremely variable and depends on several factors. Previous effective vaccination prevents or modifies the infection. In cases with high fever and in confluent and hemorrhagic types of smallpox, the prognosis is poor. The virulence of the virus in epidemics is quite variable. If complications appear, the prognosis is worse. The amount of scarring is variable but is more marked with secondary infection.

Diagnosis

The incubation period is 7 to 21 days. The prodromal illness lasts 2 to 4 days and consists of fever, extreme backache and headache, prostration and often vomiting, sore throat and cough. The onset of the eruption may be accompanied by temporary fever. The lesions are succeeded by shiny papules which become vesicles in about 3 days. On about the 8th day of eruption, pustulation occurs, followed by crusting after the 10th day. Lesions are centrifugally distributed and are most dense on the face and distal parts of the extremities. A successful vaccination usually excludes the diagnosis of smallpox in a suspected case. Infectivity is present from just before the onset until the last crust has shed.

Treatment

- A Specific Measures Hypoimmune vaccine gamma globulin
show promise experimentally
- B General Measures Penicillin has generally favorable effect
probably due to control of secondary bacterial invasion
- C Local Measures
- 1 Mouth care. Early in the disease provide good oral
hygiene (see p 5) and apply petroleum or mineral oil swab
to the nares
 - 2 Skin Care. If lesions are confluent and up-
staging treat with pyoderma (see p 84). Avoid itching by
use of antipruritics (see p 67) restraint and sedation may
be necessary
- D Treatment of Complications Treat isolated febrile
infections with appropriate symptomatic Complications
include secondary infections of the skin mucous membranes
and respiratory tract septemia nephritis myelitis and
various neurological manifestations

Prophylaxis

Vaccination (see p 494)

EPIDEMIC PAROTITIS (Mumps) (code No 621.170)

Mumps is a contagious disease caused by a specific
virus which most commonly involves the salivary glands but fre-
quently produces meningitis encephalitis orchitis and
epididymitis. The prognosis is almost always favorable. Testicul-
itis usually unilateral may follow but rarely produces sterility

Diagnosis

The incubation period is 2 to 4 weeks (usually 18 to 21 days).
Swelling of the parotid glands is the most commonest
manifestation and is usually accompanied by severe systemic mani-
festation. Headache and draw in the abdominal pain and tenderness
in the parotid swelling usually associated with fever. Generally
dendritic meningo-encephalitis (confirmed by lumbar puncture) pan-
creatitis and orchitis epididymitis. Complications including anti-
body appearance and convulsions. Mumps is probably not a
justifiable therapeutic indication of swelling and not swelling disappears

Treatment

- A Specific Measures None available
- B General Measures
- 1 Isolate until welling is gone
 - 2 Bed rest and good fluid intake
 - 3 Aspirin or non-steroidal anti-inflammatory drugs if required
 - 4 Alkaline mouth solution mouth washes
 - 5 Mumps neutral serum 20 cc mumps virus
gamma globulin 25 cc IM may reduce the incidence of
orchitis and epididymitis
- C Treatment of Complications Complications are usually
immune manifestations of the disease and treatment is
to relieve them. They may produce orchitis and epididymitis

- 1 Meningoencephalitis (code No 912 170) May be asymptomatic
 - a Analgesics as necessary
 - b Lumbar puncture if necessary to reduce headache
 - c If very severe may treat with hydrocortisone as in o chitis
- 2 Orchitis (code No 755 170)
 - a Suspension of scrotum in suspensory or toweling bridge and application of ice bags
 - b Incision of tunica may be necessary in severe cases
 - c Codeine or morphine as necessary for analgesia
 - d Infection of spermatic cord at external inguinal ring with 10 to 20 cc (2 1/2 to 5 dr) of 1% p ocal e solution
 - e Hydrocortisone 100 mg i v followed by 20 mg orally every 6 hours for 2-3 days
- 3 Prostatitis (code No 690 170) Symptomatic relief only
Parenteral fluids if necessary
- 4 Oophoritis (code No 788 170) Symptomatic treatment only

Prophylaxis

- A Mumps convalescent serum 20 cc (5 dr) i m may reduce incidence in exposed susceptible
- B Mumps virus vaccine may produce temporary active immunity
Intradermal injection of virus antigen denotes immunity if followed by local erythema

POLIOMYELITIS (Infantile Paralysis) (code No 972 171)

Acute anterior poliomyelitis is an infectious highly communicable disease caused by any one of three identified types of neurotropic virus occurring throughout most of the world in endemic form and especially in the Western world in epidemic form. It occurs with highest frequency in warm seasons. The disease is virtually as prevalent as measles but most infections are inapparent. A few produce fever and headache and back stiffness with mild transient illness. A small number (probably 0.1 to 1 per cent of cases) result in lower motor neuron palsies usually symmetrical and of great variation among patients. Case fatality rates vary according to attainment of viral quality of medical care and degree of immunity. Incidence and mortality of infection in the area. The gross fatality rate is below 5 per cent in the United States and varies from less than 5 per cent to 30 per cent in severe cases almost all of which have bulbospinal lesion of high spinal involvement with respiratory paralysis. Prognosis in severe cases depends largely upon the skill of the applier of tracheostomy and mechanical facilities for respiratory assistance and tracheobronchial toilet.

Diagnosis (For diagnosis of severe cases see p 454)

The incubation period is usually 10 to 14 days (range 3 to perhaps 30 days). The onset of paralytic disease may be preceded by a brief episode of influenza like illness followed by apparent recovery then recurrent fever headache backache neck stiffness and nausea and vomiting. Nonparalytic poliomyelitis is difficult to diagnose with certainty and is indistinguishable from a variety of

asptic meningitidis In paralytic disease weakness of paralysis may develop at any time during the febrile phase which rarely lasts longer than 7 days Cerebrospinal fluid shows pleocytosis in all but about 10 per cent of cases Differential count is not significant but the total count is high total counts with relative lymphocytosis suggest other diseases especially mumps Protein may be slightly elevated and may increase during the first few days of illness Sugar and chloride are usually normal

Virus may be isolated from the nasopharynx and stool for a few days after onset of symptoms (may be isolated as late as several weeks after first stool) Specific diagnosis is established only by isolation of the viral agent and demonstration of an increasing titer of neutralizing or complement fixing antibody in blood Serological tests are not yet widely available

Treatment (Ely Phas) (For severe cases see p 454)

A Spinal Muscles Non available

B General Measures

1 Rest and support

- Travel activity and necessary examinations and psychological should be avoided
- Profound bed rest and cursory muscle check not more than once daily in a uterus Muscular examination should not irritate muscular activity on the part of the patient
- Maintain comfortable but check positions in a padded firm mattress footboard padding rubber pads rolls saddle and light splints

2 Relieve pain and anxiety

- Aspirin or aspirin with amphetamine and phenobarbital Avoid opiates and other habituating
- Tranquillizers such as meprobamate (Equanil®), Miltown® and meprobamate (Ritalin®) may prove useful
- Hot water packs (Kneipp) to extremities or specific areas during febrile period complete body packs only when afebrile

3 Muscular Change of position extremity packs and analgesics drug usually sufficient Depot form of tubocurarine may be used but their effect has not been sufficiently evaluated

4 Bowel control Dehydration and intestinal hypomotility often lead to impaction Examine frequently Give sufficient fluids Use neostigmine and neostigmine if necessary

5 Bladder weakness May occur with paralysis involving lower extremities commonly with paraplegia

- Insert Foley catheter with greatest care
- Do not attempt to mop ophthalmia with antimicrobial
- Continue Foley catheter with gravity bottle by sterile technique Change catheter every five days and remove as soon as possible
- Test specific urinary infection after identification of organism and determination of sensitivity to antimicrobial

6 Nutrition During febrile phase and during a bedfast use nutrient diet give a maximum of 5 Gm calcium carbonate daily (or milk or milk product) and maintain fluid intake

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take to insure adequate daily output of low specific gravity urine (1.520 liters daily for adult). If nasogastric feedings are necessary, utilize liquid infant baby foods, juices, low calcium soybean milk substitutes, lactose, and vitamins.

Diagnosis (Severe Cases)

Severe poliomyelitis infections threaten life by respiratory muscle weakness, impaired airway, pharyngeal paralysis, or involvement of vital centers. Diagnosis is made upon appearance of altered breathing pattern, development of paradoxical respiration, diminution of vital capacity, pooling of pharyngeal secretions, change of voice, impairment of swallowing, diminished cough, fatigue, restlessness, lethargy, and confusion, in any combination or sequence, often in association with facial, extraocular, lingual, or masticatory muscle weakness or upper trunk—shoulder girdle paralysis. Upon the appearance of any of the manifestations of severe involvement, the following observations should be made:

- A Record predicted normal vital capacity and determine vital capacity at least daily, utilizing spirometer, basal metabolism machine, or special ventilometer.
- B Note rate of progression. Progressive involvement of muscle groups, particularly upper trunk, shoulder, and neck, with persistent fever, usually indicates fulminating course.

Treatment (Severe Cases)

A Specific Measures: None available

B General Measures

- 1 Mobilization of personnel and equipment. Symptoms of grave poliomyelitis require emergency mobilization of medical, surgical team and basic equipment, tank respirator, preferably with positive pressure attachment, tracheostomy surgical set, intravenous set (with polyethylene catheter and cut down instruments) and aspirating pump.
- 2 Indications for tracheostomy. Impaired airway from accumulated secretions, vocal cord paralysis or spasm (cyanosis, deep unconsciousness, and convulsions should not be permitted to occur), pharyngeal paralysis (impaired swallowing mechanism, regurgitation of fluid through the nose aspiration of foodstuffs), rapidly falling vital capacity, high fever, and rapid extension of paralysis. If bronchoscopic examination is performed, tracheostomy should be done with bronchoscope in situ.

Artificial respiration is usually necessary during tracheostomy, provided by means of oxygen anesthesia bag, hand respirator, or clinical respirator, e.g., positive pressure devices. NOTE: Early tracheostomy may be lifesaving and may limit extension of disease by preventing hypoxia. Indication must be met liberally, constituted with the experienced pediatric teams. With an extremely skillful nursing staff, tracheostomy may be avoided, but risk of surgery is negligible in comparison with lifesaving advantages. Use transverse incision at the level of the cricoid, with the neck extended. Insert tube through first tracheal ring below cricoid. Never perform low tracheostomy in poliomyelitis; never incise cricoid.

3 Indications for

- a Obvious respiratory and quiescent fatigue diminished expiratory volume anxiety and tachypnea
- b Vital capacity below 50 per cent of normal When vital capacity is 35 per cent of normal the patient usually is unable to cough
- c Child unable to perform coughing require expectorant clinical judgment to substitute for in the direction of safety Fatigue tachypnea disturbance of breathing pattern or lethargy warrant a trial in a respirator

4 A technique of spirometry patient with tracheostomy

- a The bronchial toilet Use separate urethral type nasopharyngeal and open ended bronchial catheters Sit the patient for maximum pressure of manometer in inches of mercury (check manometer with the bellows off!) Place the bellows in the mouth and catheter introduced the catheter with Y open closed open closed inspirations and withdraw wall Entersight in the stem of the hub by the finger held to left and valve closed Keep catheter and hand clean maintain airtight in 120 000 Zephiran® daily fresh (unbottled) 5 per cent sodium bicarbonate solution Do not dry the catheter before fitting it into the cannula
- b Rotating the Rotating respirator at least every two hours Use Teflon tubing gently Turn patient manually on the patient's side at least every half hour to prevent discomfort While tracheostomy tube is in place supply with oxygenized air oxygen stream in the cannula and the tubing open The bellows theostomy tube adapt to the tubing with plastic pipe seal cement fused
- d Obtain expectorant solution to clear the airway with the respiratory therapy Technician complex Patients are limited only to the level of the end of the tube in the trachea
- e In patients with respiratory dysrhythmias (atrial palpitations) the endotracheal intubation is a positive preliminary step with tank to maintain adequate tidal volume I deapressor catheter and diaphragm hi pressure manometer (Thoracit®) to both no depressing effect to the level of the endotracheal tube
- f For sleep apnea implant depressor guided tube possibly Do not use (methylparaffin) for tube by chloroform or in the patient Avoid bacteriostatic
- g Antimicrobial For treatment of (rarely) hemiparesis if of the patient's tracheostomy expectorant therapy Avoid administration of antimicrobial to prevent development of the tracheostomy tube
- h Maintain tidal volume and weight (See Radford et al. Nursing and the Artificial Respiration New England J Med 251:877 1954)
- i Avoid electrolyte alkalosis by control of tidal air and proper patient blood-alcohol level

C Technique of cannulation

- 1 Gargle the patient's mouth by nasogastric intubation and aspirate the epiglottis electrolyte by intravenous route

- 2 Gastric hemorrhage Uncommon, but may cause death. Transfuse if bleeding or perforated Curling's ulcer is suspected. Surgery should be undertaken under positive pressure respiration if perforation is proved.
- 3 Bladder atony and infection See page 453.
- 4 Ileus and impaction See page 453.
- 5 Atelectasia Prevent by aerosolization of air stream, periodic deep breaths by increasing tank pressure briefly or special vacuum attachment, change of position and prevention of respiratory infection as well as good tracheobronchial toilet. If atelectasis occurs treat with positive pressure aerosol, the spy bronchodilators, wetting agents and if necessary trypsin. Bronchoscopy is usually ineffective unless inspissated secretions are present.
- 6 Mental changes Psychosis (usually short lived) with confusion, disorientation and hallucinations or delusions occur in a small percentage of cases. They may be benefited by new drugs especially Frenquel® (azacyclool hydrochloride). Post acute depression is the rule in severe disease. It subsides in 6-8 weeks with supportive psychological care.
- 7 Pregnancy Pregnant women have a high susceptibility to poliomyelitis and often develop the disease. Attentive expectant care until term. If at or near term carry through labor in respirator. Deliver on detached respirator tray under positive pressure respiration with local block or do cesarean section. Mortality is negligible with a well coordinated respiratory and obstetric program. Early in pregnancy spontaneous abortion may occur or surgical abortion may be necessary. Try to avoid procedure until the end of the febrile period.

D Convalescence and Rehabilitation

- 1 Principles Prevent deformity, avoid exercise during febrile period and mobilize early range of motion and position change during febrile period. Early active exercise under skilled direction as soon as feasible.
- 2 Early bracing and splinting for therapeutic purposes in order to activate therapy program. NOTE: The full gamut of physical and occupational therapy, individual and group psychology, social service and application of all medical specialties may be required in the rehabilitation process.

Prophylaxis

A safe and effective killed virus vaccine which is antigenic for all three types of virus is now available and officially recommended for people so as through age 40. It can be given to infants at the same time as DPT or pertussis immunization. For children or adults give 1 cc intramuscularly after two to four weeks and 1 cc after a venereal infection. A second injection will probably prove to be necessary.

Immune reaction does not prevent survival of the virus if the infectious agent and outbreaks may still be anticipated in crowded areas. During such outbreaks, avoid fatigue and contact with crowds should be minimized.

The incidence of other neurotropic virus infections warrants a regime of rest and close observation. Fall ill divided the people with children with symptoms of meningeal irritation.

PSITTACOSIS (Ornithosis) (code No 010 173)

Psittacosis (ornithosis) is characterized by pneumonitis often migratory usually associated with fever to emia. A history of contact with parrot, parakeets, pigeons or rarely other birds is usually obtainable. Diagnosis is proved by isolation of virus from blood or sputum of the patient or by rising titer of complement fixing antibody. Human to human transfer is rare although isolation precautions are wise.

Treatment consists of giving tetracycline drugs or chloramphenicol 0.5 Gm. every 6 hours orally or 0.5 Gm. I.V. every 12 hours for 10-14 days or penicillin (aqueous) 100,000 units I.M. every 3 hours for 2 weeks. Give oxygen and sedation as required.

ENCEPHALITIS (code No 930 1)

Encephalitis may be athropod born (Eastern and Western equine encephalomyelitis, St. Louis or Japanese B) post infective (measles, vaccinia, etc.) or of unknown type (von Economo's et al.). Severe and protracted neurological abnormalities signs of meningeal irritation and convulsions may be noted. Lymphocytosis is found in the cerebrospinal fluid unassociated with decrease of a gamma globulin level. Rising titer of complement fixing neutralizing antibody is confirmatory in the atropod born types and in mumps encephalomyelitis.

Repeated lumbar punctures may relieve symptoms. Prevention of alytic treatment of diseases, pneumonia and urinary tract infection is important. Give anticonvulsants as needed (pp 351 and 533).

LYMPHOCYTIC CHORIOMENINGITIS (code No 910 160)

Lymphocytic choriomeningitis is a viral infection of the central nervous system which is clinically undistinguishable from non-polytrophic poliomyelitis or mild encephalitis. The virus may be isolated from the blood or spinal fluid of diagnosis may be confirmed by rising titer of neutralizing or complement fixing antibody.

Incubation period approximately 8 to 21 days.

Treatment for encephalitis (see above).

DENGUE (code No 010 162)

Dengue is an acute infectious disease caused by a virus transmitted by mosquitoes. The incubation period usually 5 to 8 days following the bite of an infected Aedes mosquito. Onset with illness is aching of the back and extremities, chills and fever prostration. Conjunctival injection and generalized lymphadenopathy may be found. The fever usually lasts 5 to 6 days and may be of the saddle back form. Aseptic meningitis or meningoencephalitis may occur on the third to fifth day and last up to 3 days. Leukopenia is usually marked. Fatality is extremely rare.

Give salicylates as required for discomfort. Permit gradual restoration of activity during prolonged convalescence.

Available prophylactic measures include control of mosquitoes by screening and DDT. Dengue vaccine shows promise experimentally.

RABIES (code No 010 174)

Rabies is an acute viral infection primarily of animals which is occasionally transmitted to man. It is characterized by apathy or hyperexcitability, paralysis, and invariably results in death.

Diagnosis

The incubation period is usually 2 to 8 weeks (occasionally as long as 1 year) following the bite of a rabid animal. Onset occurs with pain and numbness at the site of inoculation followed by depression, irritability, and mild dysphagia. This is followed by hyperesthesia and muscle spasms, particularly of the pharynx. Eventually paralysis and death occur.

Treatment

Treatment consists of absolute quiet and freedom from stimulation and sedation as in tetanus to prevent triggering convulsions. No specific measures are available.

Prophylaxis

- A Observation of animal producing bite
- B Cauterization of wound with fuming nitric acid followed by neutralization of the acid with lime water or thorough washing with green soap
- C Rabies vaccine 2 cc subcutaneously for 14 days following positive diagnosis of rabies or following bite by a suspected animal if animal cannot be observed or found to be on the head
- D Rabies hyperimmune serum should be administered in addition to vaccine in fatal or severe hand bites

YELLOW FEVER (code No 010 178)

Yellow fever is a fatal infection of man and monkeys due to a virus transmitted by Aedes mosquitoes. It is characterized by fever, relative bradycardia, and hemorrhagic phenomena. The mortality rate of yellow fever is quite variable including missed cases; it is probably 5 percent.

The incubation period is 3 to 6 days. The onset is with chills, headache, and backache. The fever often is *diurnal* after 3 days during which time the patient usually sheds a toxic and may have severe nausea and vomiting. The conjunctiva are injected and the tongue red. This is followed by pallor, chills, bleeding gums, black vomit, slight jaundice, melena, albuminuria, and prostration. Leukopenia and relative bradycardia are usually seen.

Treatment consists of giving a liquid diet, limiting food to high carbohydrate, high protein liquids as tolerated, intravenous glucose and saline as required, analgesics and sedatives as required, and saline enemas for obstipation.

Available prophylactic measures in the mosquito control by adequate screening DDT etc. Vectors are available. Give 0.5 cc b.c.u.

INFLUENZA (code No 010 168)

Influenza is a acute viral infection of the respiratory tract characterized by abrupt onset of systemic and respiratory symptoms. The antigenic strains of influenza virus exist (A B C). Most preventable infectious disease.

In treatment bed rest to avoid complications. The most important consideration is the isolation and adequate hygiene. May be a self-limiting disease of only a few days. Prophylactically a dose should be given for treatment of bacterial complications such as pneumonia.

Polyvalent influenza vaccine is the most effective. One should be given but highly effective against influenza epidemics. Usually it is built up by season.

DISEASES DUE TO RICKETTSIAE

The rickettsia is a arthropod born organism which spread by the blood of the host. The virulence of the rickettsia varies with the different species. The rickettsia of the typhus group is a epidemic typhus. Rocky Mountain spotted fever and typhus are the most common. The advantage of the rickettsia is that it is highly infectious.

D. gnosis

A. Typhus fever (code No 010 184). Incubation period 6 to 15 days. Typhus fever is caused by Rickettsia prowazekii (p. dem type) or Rickettsia mooseri (mu. typ). The former transmitted by body louse the latter by rat fleas. The disease is similar except that the mortality is less. Onset abrupt with high fever, aching and prostration. Delirium, toxicomania, myoclonus. A macular papular rash begins on the fourth to seventh day. Protrusion of the tongue and spreading to the extremities usually spreading to the palms and soles. Leukopenia. Usually diagnosis may be confirmed by complement fixation. Positive OX 19 agglutination appears during the disease.

B. Rocky Mountain spotted fever (code No 010 181). The incubation period 3 to 14 days. Rocky Mountain spotted fever is caused by Rickettsia rickettsii and transmitted by tick bite (p. inc. pally. De m. t. and om. and D. variabil.). Prodromal symptoms of malaise, noxa and malaise may occur. The onset of high fever, headache, photophobia, pain in the muscles usually a dose. Arterial and renal hemorrhagic maculopapular rash appears first on the wrist and ankle and then spreads to the face and palms and

soles. Leukopenia is usually present early. Complement fixing antibodies and agglutinins for *Proteus* OX 2 or OX 19 appear during the second week.

- C Scrub Typhus (Tsetsu g mushi Fever) (code No 010-183) The incubation period is 6 to 18 days. Scrub typhus is caused by *Rickettsia orientalis* (*R. tsutsugamushi*) and is transmitted by larval mites. An eschar at the area of inoculation is common. The onset is sudden with chills, fever, malaise, and cough. A dull red maculopapular eruption appears on the trunk from the fifth to eighth days and may extend to the extremities. Specific complement fixing antibodies or *Proteus* OX K agglutinins appear during the second week.

- D Q Fever (code No 010-185) The incubation period is 14 to 26 days. Q fever is caused by *Coxiella burnetii* and is apparently acquired from sheep, goats, and cattle in a manner not yet determined. Headache, fever, cough, and stiff neck are common symptoms. Repeated rigors may occur. Pneumonitis or hepatitis may be demonstrated. Specific complement fixing antibodies appear during the second or third weeks.

- E Rickettsialpox (code No 010-187) *Rickettsialpox* is an infection caused by *Rickettsia akari* introduced by the bite of a mite. A lesion which passes through the stage of papule, vesicle, and eschar precedes the onset of fever, chills, headache, photophobia, and muscular aches by about a week. A generalized rash which evolves through papular, vesicular, and crusting stages appears at the onset of fever or a few days later. Leukopenia usually is present.

Treatment

A. Specific Measures

1. Tetracycline 0.5 to 1.0 Gm orally every 6 hours for 2 to 7 days or 0.5 Gm I.V. every 12 hours.
- or 2. Chloramphenicol U.S.P. (*Chloromycetin*®) 0.5 Gm orally every 6 hours for 2 to 7 days.

B. General Measures

1. Parenteral fluids, oxygen, and sedation as required.
2. Other supportive measures as needed.
3. Decontaminating procedures must be carried out for louse-borne infections (see p. 92).

Prophylaxis

A. Specific Measures

1. *Typhus* (Cox type) 1 cc subcutaneous at 7 to 10 day intervals.
2. Rocky Mountain spotted fever vaccine 1.0 cc about 3 times at 5 to 7 day intervals.

- B. General Measures. Decontaminating is very important in louse-borne epidemic typhus (see p. 92).

DISEASES DUE TO BACTERIA

SCARLET FEVER (code No 010 102) and
STREPTOCOCCIC SORE THROAT (code No 531 102)

Scarlet fever and streptococci sore throat (follicular tonsillitis and suppurative throat) are infections of the fauces by *β* hemolytic streptococci (Lancefield Group A). In scarlet fever in addition to the infection manifested by the organism toxin a prominent feature is the rash.

The mortality rate from streptococci sore throat and scarlet fever in the United States is 0.5% or less. Rheumatic fever is probably more common than generally supposed and may be apparent only when serial electrocardiograms are taken during convalescence.

Diagnosis

The incubation period is 2 to 7 days. The onset is usually abrupt with chills, fever, headache, pain in the extremities and swollen lymphatic nodes and sore throat. The throat is usually follicular and moderately edematous. Exudate appears as concentric patches of whitish material which may be easily wiped off. The rash of scarlet fever consists of a punctate erythema which is most distinct on the skin fold of the axilla and groin but does not appear on the extension of the surface of the upper extremity. Strawberry tongue and stippling of the soft palate may be noted. Diagnosis may be confirmed by culture. Sedimentation rate is raised leukocytosis is present.

The duration of the infection varies. It may be prolonged by a convalescent period.

TreatmentA. Specific Measures

1. Penicillin procaine 300,000 units daily I.M. Penicillin may be continued 5 to 7 days. Penicillin may be given orally. Oral penicillin 200,000 units every 6 hours. Benathin Penicillin G U.S.P. (Bicillin®) 1,200,000 units I.M. may be used. Local penicillin by lozenges with saline.
2. Tetracycline drugs 0.25 to 0.5 Gm. every 6 hours or erythromycin 0.2 to 0.5 Gm. every 6 hours is effective but may be followed by bacterial dysbiosis and clinical relapse.
3. Sulfonamide have no effect on the course of streptococci sore throat. If given for 2 weeks. Dose 0.5 Gm. (7½ gr.) every 4 hours with equal or double quantities of sodium bicarbonate.
4. Scarlet fever streptococci Antitoxin (8,000 to 36,000 units) may be given I.M. with benefit in severe toxic cases of scarlet fever.
5. Convalescent serum 25 to 150 cc. (1.5 oz.) may be used similarly to tetracycline and may be given I.V.

B. General Measures

1. Bed rest until all bruits and diminution in rate is normal.
2. Do not attempt to suppress the fever.
3. Hot saline or 30% glucose gargles or throat irrigation 3 or 4 times daily.

4 times daily for relief of sore throat

4 Aspirin or codeine as necessary for symptomatic relief

C Treatment of Complications

1 Complications due to infection include cervicofacial adenitis, rhinitis, otitis, otitis, mastoiditis, pneumonia, empyema, septic arthritis and septicemia. Treatment with penicillin is usually effective (see p 502).

2 Complications of unknown etiology

a Rheumatic fever may be prevented by early vigorous treatment of the infection with penicillin (see p 518).

b Acute hemorrhagic glomerulonephritis (see p 293).

D Treatment of Carriers 300 000 units of penicillin procaine complex daily I M for 10 days or benzathine penicillin G 1 200 000 units I M usually abolishes the carrier state.

Prophylaxis

A Scarlet fever toxin in 5 weekly injections of 500 2000 8000 25 000 and 80 000 units because it prevents the toxic manifestations of scarlet fever but does not prevent streptococcal infection.

B Sulfonamides 0.5 Gm (7½ gr) b.i.d. penicillin 100 000 units by mouth b.i.d. or benzathine penicillin 600 000 units I M once a month recommended in case of streptococcal infection. These should be reserved for individuals with rheumatic lesions to prevent recurrence of rheumatic fever.

DIPHTHERIA (Pharyngeal code No 631 125)

(Laryngeal code No 330 125)

(Nasopharyngeal code No 318 125)

Diphtheria is an acute infectious disease caused by *Corynebacterium diphtheriae* and is characterized by the formation of a pseudomembrane at the portal of entry usually the respiratory tract and by the activity of exotoxin at distant sites.

The mortality rate of diphtheria generally varies between 10 and 20 percent. Older individuals do poorly and delay of treatment carries with it a high mortality rate. Myocarditis appears early and is frequently fatal and disturbances of conduction or the appearance of rapid arrhythmias imply lethally prognosis. However, if the patient survives recovery is usually complete. Neuritis is rarely fatal and permanent complication paralysis of cranial nerves or nuchal rigidity is usually fatal. Surviving patients recover slowly but completely.

Diagnosis

The incubation period is 2 to 7 days. Symptoms develop at the site of the lesion but include sore throat, nasal discharge, hoarseness accompanied by malaise and low grade fever. The pseudomembrane is typically grayish homogeneous and dense. Edema and a narrow zone of erythema around the lesion usually found. Diagnosis is confirmed by culture. If culture is negative assumed as long as *C. diphtheriae* present in the nasopharynx the carrier state is not uncommon.

TreatmentA Specific Measures

- 1 Diphtheria antitoxin must be given in all cases where diphtheria cannot be excluded by simple clinical examination. The intravenous route preferable in all except the mild cases in those who are unfit to tolerate a rumen Conjunctival and conjunctival for a rumen sensitivity (see p. 495) should be given in all cases and desensitization (see p. 495) is recommended. The dose varies with the duration of the disease, the location of the lesion and the age of the patient. A single dose should suffice.

Diphtheria Antitoxin Dosage Schedule

Location	Child	Adult
Anterior nasal	5000 units	10,000 units
Mild pharyngeal	10,000 units	20,000 units
Moderate pharyngeal	20,000 units	40,000 units
Severe pharyngeal and nasopharyngeal	40,000 units	80,000 units
Laryngeal	10,000 units	20,000 units
Any two sites	40,000 units	80,000 units
Late case		

- 2 Penicillin procaine 300,000 units daily. Penicillin 50,000 units every 3 hours. If reaction slightly the dose per patient with organism from the throat and cuts again. A dry throat, clinical evidence of disease not alter the course of the diphtheria is left.

B General Measures

- 1 Absolute bed rest for at least 3 weeks and until ESR is normal.
- 2 Liquid diet diet is tolerated.
- 3 Hot saline or 30% glucose throat irrigation 3 or 4 times daily.
- 4 Aspirin or codeine as required for pain.

C Treatment of Complications

- 1 Myocarditis (code No. 430.125.9). This may occur at any time post several weeks after onset and may be associated with peripheral vascular collapse. Anginal or abdominal pain, nausea and vomiting, cyanosis may be noted. Detection of the mitral regurgitation, second degree in blood pressure, gallop rhythm or any arrhythmia may be found. ECG evidence is usually demonstrated in several records.
 - a No definitive treatment known.
 - b Oxygen by tent or mask may be needed. Hypertonic glucose solution 100% of 20% solution daily may aid.
 - c Digitalis and quinidine should be reserved for paroxysmal arrhythmias.
- 2 Nerve (code No. 98.125.9) generally develops at beginning of the 3 weeks after the onset. Nerve degeneration of fluid through the nerves (NIX) paralysis of accommodation (NIII) dysphagia and dysphonia (NX) and rarely involvement of other cranial nerves usually preceded involvement of

the child emits which is associated with paroxysms weakness and depression of reflexes. Nasal feeding should be attempted in such cases. Corrective splinting and physical therapy may be of aid.

3 Respiratory tract obstruction. Croupy cough, stridor and dyspnea, great laryngeal obstruction.

a Suction of membrane and secretions under direct laryngoscopy may help.

b Intubation or tracheotomy should be performed before the appearance of cyanosis if the distress increases.

D Treatment of Carrier. Penicillin has very limited effect on the carrier state.

Prophylaxis

A Children. Three injections (0.5, 1.0 and 1.0 cc) of diphtheria toxoid (formol alum or aluminum hydroxide precipitated) at one month intervals during the second 6 months of life (may be combined with tetanus toxoid and pertussis vaccines). Follow by Schick test in 3 to 6 months. Give 1.0 cc booster at 2 years and at start of school.

B Adults

- 1 Sensitivity test. Moloney's test for sensitivity to toxoid. 0.1 cc of 1:20 dilution of plain toxoid intradermally. Read like Schick test at 24 to 48 hours.
- 2 If Moloney test is negative proceed as in children.
- 3 If Moloney test is positive give 0.1 cc of 1:10 dilution of formol toxoid intramuscularly at 3 week intervals for 3 doses.

PERTUSSIS (Whooping Cough) (code No. 350.108)

Pertussis is an acute communicable infection of the respiratory tract. It is caused by *H. pertussis* and is characterized by paroxysmal cough, whoop, and absolute lymphocytosis. Until recently the mortality rate of pertussis in infants under one year of age with disease is most common was 20 percent. This has been materially reduced with modern antibacterial therapy. Older children rarely die of pertussis.

Diagnosis

The incubation period is 7 to 14 days. The onset is with coryza followed by gradually increasing cough. After 1 to 2 weeks the cough becomes paroxysmal especially at night and is often followed by vomiting. Whoops may be heard during the paroxysm but are often absent or infrequent in young infants. Absolute lymphocytosis appears during the paroxysmal stage. The diagnosis may be confirmed by cough plate, nasopharyngeal swab cultured on Bordet-Gengou's medium. Infectiousness greatest early in the disease and decreases until the organisms disappear from the nasopharynx after about one month.

Treatment

A Specific Measures

- 1 Antibiotic

- a Tetracycline 25-50 mg/Kg (11 mg/lb) per day orally
- or b Streptomycin 1.0 Gm per day in divided doses I.M. for one week may be effective
- c Chloramphenicol USP (Chloromycin®) 50 mg/Kg (23 mg/lb) per day orally
- d Polymyxin 2.0 mg/Kg (1 mg/lb) per day I.M. also promising but this drug may be toxic (see p 510)
- or e Erythromycin 30 mg/Kg/day orally
- 2 Hypersensitive serum hyp immune gamma globulin appears to be ten percent prevent complications and reduce mortality 20% hyp immune serum or 2.5% hyper immune gamma globulin given daily or every other day I.M. for 4 or 5 doses

B General Management

1 Nutrition

- Frequency of feeding may be necessary
- b Re-feed if vomiting occurs shortly after a meal
- High caloric formula by gavage tube may be required in infants who refuse to eat
- d Parenteral fluids may be used to insure adequate fluid intake as necessary

2 Cough

- a Sedative and expectorant cough mixture as a rule of only light benefit
- b Atropine titrated to the point of futility by increasing doses of tincture of belladonna every 4 hours starting with one drop once a day helps
- Ethereol by rectum may be desirable

C Treatment of Complications

- 1 Cerebral hemorrhage talp apnea and umbilical hernia may occur due to the increased pressure caused by the cough. Treat symptomatically
- 2 Pneumonia usually develops secondary invade should be treated by hyp immune gamma globulin (see above) penicillin and sulfonamides or tracheostomy in Oxygen is often required
- 3 Convulsions may be due to a toxemia petechial hemorrhage the brain substance encephalitis or cerebral hemorrhage If convulsion resistant 100% oxygen inhalation and lumbar puncture may be of aid
- 4 Otitis media should be treated with penicillin or sulfonamides Myringotomy may be necessary

Pertussis

- A Passive prophylaxis of exposed child may usually be accomplished by the injection of 20 cc of hyp immune serum 2.5 cc of hyp immune gamma globulin I.M.
- B Active immunity may be produced by the subcutaneous injection of a vaccine containing 10 billion organisms per cc. The initial dose consists of 1 cc and this is followed at 2 to 4 weeks in intervals by the second and third doses of 2 cc each. Vaccine containing 30 billion organisms per cc is also used. Follow the manufacturer's instructions on the package. Immunity is attained at 6 months of age with usual procedure but immunity may start at 2 months if properly followed. Booster shots of

1 cc are given at 2 years and when starting school. The vaccine may be combined with diphtheria and tetanus toxoid. Encephalitis occasionally follows pertussis immunization.

MENINGITIS

MENINGOCOCCIC (Epidemic) MENINGITIS (code No 910 104)

Epidemic meningitis caused by *Neisseria meningitidis* follows a bacteremia from a nasopharyngeal focus localizing in the meninges. Bacteremia acute or chronic may occur without meningitis. The overall mortality of epidemic meningitis is 10 per cent. Young healthy individuals and those who retain consciousness usually survive.

Diagnosis

The incubation period is 3 to 7 days. Fever, chills, headache, pain in the back, abdomen and extremities, and nausea and vomiting may be present. Delirium, stupor, or coma appear in severe cases. A petechial rash is commonly seen. The neck is stiff and Kernig's and Brudzinski's signs are positive.

Lumbar puncture reveals a cloudy spinal fluid under increased pressure usually containing more than 1000 cells per cubic mm with polymorphonuclears predominating. The spinal fluid glucose and chloride contents are decreased and the protein increased. The organism is usually demonstrated by smear or culture.

Infectivity may be present for a few days before the appearance of meningeal invasion but is quickly terminated by therapy. Case to case spread is uncommon. Healthy nasopharyngeal carriers are the common source of spread.

Treatment

A. Specific Measures

1. Sulfonamides The agent of choice

- a. Severe cases. Sodium sulfadiazine or diaminodiphenyl sulfamide or a mixture of equal parts of each or sulfisoxazole. Give 5.0 Gm (75 gr) in 1000 cc (2 pts) of an electrolyte solution preferably Ringer's lactate solution I.V. or subcut by electrolysis.
- b. Mild cases. Sulfadiazine, sulfamethoxazole or a mixture of equal parts of each or sulfisoxazole. Give 3.0 Gm (45 gr) orally with equal or double amount of sodium bicarbonate.
- c. Follow by 3.0 Gm (45 gr) I.V. or subcut every 8 to 12 hours or 1.0 Gm (15 gr) orally every 4 to 6 hours as indicated by severity. Give equal or double doses of sodium bicarbonate orally with the sulfonamide.

and 2. Penicillin

- a. Aqueous penicillin 100,000 unit I.M. every 3 hours
- or b. Penicillin procaine 600,000 units I.M. twice daily

3. Antibacterial therapy need only be continued on weeks.

B. General Measures

1. Sedation with paraldehyde and meprobamate I.V. or meprobamate sulfate as necessary for restlessness.

- 2 Restrict intake if necessary for marked r stlessness
- 3 Fluid intake should be at least 3000 cc (6 pts) daily and should be sufficient to maintain a urinary output of at least 1000 to 1500 cc (2 3 pt) . Replace fluid lost by vomiting and give paracetamol orally as needed
- 4 Feedings (and medication) by stomach tube if comatose more than 3 days
- 5 Lumbar puncture to be repeated if evidence of increased intracranial pressure persists or to be kept responsive to therapy by cerebrospinal fluid glucose level
- 6 SHOCK Treat for shock (p 27) and for a uterine adrena insufficiency (p 38)
- C Complications and Treatment Myocarditis and renal failure are common in severe cases as is no response if treatment is available Cranial nerve lesion (especially N VIII) are usually permanent Arthritis requires no treatment other than paracetamol for pain

Pneumonia

A total of 1.0-2.0 Gm (15-30 gr) sulfadiazine orally to exposed persons or carriers in two doses taken the same day

PNEUMOCOCCIC MENINGITIS (code No 910 101) STREPTOCOCCIC MENINGITIS (code No 910 102) STAPHYLOCOCCIC MENINGITIS (code No 910 105)

Symptoms are similar to those of meningococcal meningitis but spreading infection usually present and a focus is often demonstrable in the lungs (pneumonia) the middle ear or sinus . The spinal fluid must be obtained and examined to determine the causative agent

Treatment

A Specific Measures

- 1 Treat as meningococcal meningitis (p 46) and also give 10 000 units of penicillin in 10 physiological saline intrathecally once daily until CSF glucose normal
- 2 Add aqueous penicillin 1 000 000 units 1 M e very 2 hours
- 3 Erythromycin 0.5 Gm every 6 hours for staphylococcal meningitis
- 4 Tracey line drugs may also be effective do not combine with penicillin

B General Measures Symptomatic and supportive

HEMOPHILUS INFLUENZAE MENINGITIS (code No 910 110)

The form of meningitis may develop gradually suddenly usually occurs in infant less than 2 years old The symptoms are similar to those of any purulent meningitis (p 466)

Treatment

A Specific Measures

- 1 Dihydrostreptomycin (adults 1.0 Gm child 250 mg) 1 M e every 6 hours for one week and streptomycin 35 mg in 10 cc physiological saline intrathecally daily

468 Typhoid Fever

- until cerebrospinal fluid glucose content is normal
- 2 Severe cases should also be given a Madiazine sulfamerazine or a mixture of equal parts of each 150 mg /Kg body wt (65 mg or 1 gr /lb) per day Give equal or double doses of sodium bicarbonate
 - 3 Hemophilus influenzae antiserum (Type B) should also be given to patients with delayed response 25 to 100 mg of antibody nitrogen daily I V until serum produces quelling reaction
 - 4 Tetracycline drugs 0.5 Gm every 6 hours are of value Chloramphenicol (Chloromycetin[®]) is also effective
- B General Measures** Treat symptoms as they arise and maintain good nutrition and adequate fluids

TUBERCULOUS MENINGITIS

(Leptomeningitis code No 912 123)
(Pachymeningitis code No 911 123)

This disease is caused by spread of the tubercle bacilli from a focus usually in the lungs or in the peritracheal peribronchial or mesenteric lymph node There is usually a gradual onset of symptoms with listlessness irritability anorexia and fever followed by headache vomiting night cries convulsions and rigidity of the neck opisthotonos paralysis and coma

Cerebrospinal fluid frequently xanthochromic under increased pressure and on standing may form a web and pellicle from which organisms may be demonstrated by smear culture or guinea pig inoculation Cells and protein increased but sugar and chlorid are decreased

Treatment

A Specific Measures

- 1 Streptomycin dihydrostreptomycin 30 mg /Kg body wt of each per day I M in divided dose every 8 to 12 hours for 5 months Streptomycin 2 mg /Kg body wt intracally daily for 2 weeks every other day for 2 weeks and twice a week for 2 weeks (intracally therapy probably unnecessary if isoniazid used)
- and 2 Isoniazid (INH) 10 mg /Kg body wt per day divided into 2 or 4 doses for one year
- or 3 Aminosilylic acid (PAS) 3 to 5 Gm every 6 hours by mouth or sodium paraaminosalicylate 15 to 30 Gm daily I V for one year

B General Measures Treat symptoms as they arise and maintain good nutrition and adequate fluids

TYPHOID FEVER (code No 010 115)

Typhoid fever is an infectious disease caused by Eberthella typhosa a characteristic by bacterial ulceration of the lymphoid tissue of the small intestine and generalized toxemia The mortality of typhoid fever varies from 5 to 25% Elderly individuals do poorly

Diagnosis

Incubation period is 5 to 14 days. A gradual onset of fever and malaise often associated with cough or epistaxis is followed by a period of 2 or more weeks of sustained fever and then gradual defervescence. Rose spots, splenomegaly, relative bradycardia, dulcified pulse and pea soup stool may be noted. Leukopenia is the rule. Isolation of the organism from the blood, stool or urine or a high or rising titer of agglutinin confirms the diagnosis.

Infectivity begins with the appearance of E. typhosa in the stool or urine and continues until the organism disappears from the excreta which occur at varying times later. Carriers can be a serious public health problem.

TreatmentA. Supportive Measures

1. Chloromphenicol U.S.P. (Chlormoxylin®) 1.0 Gm every 4 hours by mouth until afebrile and then 0.5 Gm every 6 hours. (Children: 50 mg/Kg body wt./day followed by 25 mg/Kg/day when able to take by mouth) Continue treatment for 3 weeks.
2. Hydrates: 20 mg every 6 hours may be used temporarily in severely toxic patients.

B. General Measures

1. Prevention of dehydration by careful bathing, skin massage and use of rubbered gloves over palms and soles.
2. Careful oral hygiene.
3. Intake of food and liquids is important.
 - a. High caloric liquid diet (approximately 3600-4800 Calorie per day) (See diets pp. 55-57). Complete vitamin supplementation must be added.
 - b. Caloric diet: About 1500 Calories/liter (1420/qr)

Lactose	400 Gm	8 oz
Corn	800 cc	12 z
Milk	2800 cc	6 pts
 - c. Cassin hydrolysat formula: about 1050 Calories/liter (1016/qr)

Cassin hydrolysat	125 Gm	4 z
Milk	1000 cc	2 pts
4. Adequate fluids: Parenteral glucose solution may be necessary if oral fluid intake is diminished in unconscious patient.
5. Distension may be relieved by gentle colonic flaccid abdominal massage. Pilocarpine® and castor oil may be used with great caution because of danger of perforation.
6. Diarrhea may be controlled by tincture of opium (see p. 259).
7. The patient must be strictly isolated and his excreta sterilized until negative stool cultures have been obtained.

C. Treatment of Complications

1. Pneumonia (pneumonia) may be treated with penicillin, sulfamid, streptomycin, or tetracycline drugs depending on the etiologic agent (see p. 514).
2. Hemorrhage is usually manifested by icterus, purpura and drop in blood pressure associated with the appearance of gross blood in the stool. Transfusion should be used as required.

3 Perforation is often accompanied by abdominal pain, shock and signs of peritoneal irritation. Immediate surgery is required, anticipate and treat shock (see p. 27) before it is manifest.

D Treatment of Carriers Chemotherapy is usually ineffective in abolishing the carrier state.

Prophylaxis

A Typhoid vaccine (1 billion organisms per cc) 0.5 cc, 1 cc and 1 cc at weekly intervals by subcutaneous injection. This is usually given with paratyphoid A and B vaccine. Intradermal injection of 0.1 cc at weekly intervals may be used to minimize unfavorable reactions.

B Drinking water and milk must be boiled during an epidemic.

C Carriers must be rigidly controlled and not permitted to be food handlers.

PARATYPHOID FEVER (code No. 010.114)

This disease may be caused by either *Salmonella paratyphi* (paratyphoid A) or *Salmonella schottmulleri* (paratyphoid B) and is clinically similar to typhoid fever. Paratyphoid fever, however, is usually more abrupt in its onset and usually is a milder disease than typhoid. The differential between these infections can only be made by bacteriological examination.

Treatment and prophylaxis are as for paratyphoid fever.

BRUCELLOSIS (Undulant Fever)

Brucellosis is an acute or chronic systemic infection due to *Brucella melitensis* (code No. 010.1171), *Brucella abortus* (code No. 010.1172) or *Brucella suis* (code No. 010.1173) and is usually acquired by the ingestion of infected dairy products or by contact with infected material.

Death is not common in brucellosis but may occur from prolonged debility or subacute bacterial endocarditis. The acute stage may terminate spontaneously or may become chronic. The chronic form may persist for many years.

Diagnosis

A history of ingestion of raw milk or butchering of infected animals is helpful. Human to human spread does not occur. Fever may be a periodic undulating low grade or absent for long periods. Systemic symptoms include malaise, arthralgia, depression and sweats. Splenomegaly may be present. Leukopenia is the rule. Positive blood culture is diagnostic. High or rising titers of agglutinating complement fixing antibody in the presence of a compatible clinical picture allow diagnosis. Skin tests and opsoninophagocytic index are unreliable evidence of active infection and the former may induce confusing body responses.

Treatment

A Spectro-Tetracycline The effects of tetracycline drugs
chloramphenicol (Chloromycetin®) and di-
phenylhydantoin (Dilantin®) in

sulfadiazine therapy is not based entirely established in chonic buellosis

- 1 Combination of tetracycline 2.0 Gm daily and tetracycline drugs 2.0 Gm daily is probably the treatment of chills
- or 2 Tetracycline drug orally Give 50 mg on the first day 50 mg twice the second day 50 mg 3 times the third day a dose of 1.0 Gm every 8 hours for the following 12 to 14 days Small initial dose avoids Herxheimer like reaction
- or 3 Chloramphenicol U.S.P. (Chloromycetin[®]) 60 mg/Kg body wt (27 mg/lb) orally initially then 0.25 Gm every 3 hours until febrile 7 days
- 4 Dihydrostreptomycin 0.5 Gm I.M. every 8 hours for 2 weeks and sulfadiazine 1.5 Gm (45 g) initially and 1.0 Gm (15 g) every 6 hours for 2 to 3 weeks

B General Measures

- 1 Bed rest in a comfortable stage
- 2 High vitamin intake

Prophylaxis

- A Destruction of infected dairy animals and immunization of susceptible animals
- B Pasteurization of all milk and milk products

GAS GANGRENE (code No 12)

Gangrene is a type of severe necrotic Clostridial infection complicating infection of wounds which are soiled with dirt or feces. Destruction of tissue is usually present. Fever, chills and local swelling are usually present. Gas bubbles in the tissue may be demonstrated by x-rays. Bacteremia may be present. Anaerobic culture of discharge from the wound should confirm the diagnosis.

Treatment

- A Specific Measures
 - 1 Penicillin 100,000 unit I.M. every 3 hours
 - and 2 Polyvalent gas gangrene antitoxin 20,000 unit at once and repeat every 8 to 12 hours
 - and 3 Full doses of sulfadiazine sulfamethoxazole mixture (see p 499)
- B Supportive Measures
 - 1 Adequate fluid intake and excretion
 - 2 Infected

TETANUS (code No 010 11x)

Tetanus is a nervous system intoxication caused by fixation of Clostridium tetani toxin which enters through an open wound and infection is irreversible. The disease is characterized by tetanic contractions of striated muscle.

Diagnosis

period is 5 days to 5 weeks. The first symptoms

toms usually are pain and tingling at the site of inoculation which may be an insignificant wound that has become contaminated with soil. This is followed by irritability, trismus, stiffness and extremities and spasm of the abdominal muscles. Rigidity of muscles, risus sardonicus, stiff neck, rigid abdominal muscles and hyperactive reflexes are found. Tonic convulsions gradually appear and are precipitated by slight stimulation. Fever is variable. Death usually occurs from asphyxia or pneumonia. The mortality rate of tetanus is approximately 40%. A long incubation period and the delayed appearance of tetanic convulsions are favorable signs.

Treatment

- A Specific Measures Tetanus antitoxin 60 000 units I V
- B General Measures
 - 1 Absolute quiet with minimum stimulation
 - 2 Anti convulsant drugs
 - a Thiopental U S P 15 to 25 mg /Kg body wt ($\frac{1}{16}$ gr /lb) rectally every 1 to 4 hours as required to prevent convulsions
 - or b Sodium Amytal® 5 mg /Kg of body wt ($\frac{1}{32}$ gr /lb) I M as required
 - or c Paraldehyde 4 to 8 cc orally or 10 to 40 cc rectally p r n
 - 3 Curare in oil and beeswax shows experimental promise but is not yet established
 - 4 Mephenesin (Tolserol®) 1 to 3 Gm orally or 1 to 2 Gm I V (2 to 5% solution) may be combined with barbiturates
 - 5 Intravenous fluids as required

Prophylaxis

- A Tetanus toxoid 1 cc in 3 doses at 3 to 4 week intervals followed by booster of 1 cc at one year and 1 cc at time of injury
- B Tetanus antitoxin 6000 units I M in individuals having soil contaminated wounds especially puncture wounds compound fractures and powder burns. Adequate debridement
- C Adequate debridement of wounds

BOTULISM (code No 010 120)

Botulism is a food poisoning caused by the ingestion of preformed toxin of *Clostridium botulinum* and characterized by involvement of the central nervous system especially the bulbar region. The mortality of botulism is 60 per cent depending on the time from onset to 10 days.

Diagnosis

Symptoms appear 18 to 36 hours after the ingestion of improperly sterilized foodstuffs usually home canned. Weakness, vertigo, ptosis, strabismus, dysphagia and dysphonia are usually noted. Gastrointestinal symptoms are generally slight or absent and fever is not prominent. Usually several members of a family are involved. Death occurs from respiratory failure or pneumonia.

Treatment

A Specific Measures Botulinus antitoxin Types A and B
10 000-50 000 units of M as soon as possible

B General Measures

- 1 Absolute rest with foot of bed elevated to promote drainage from respiratory tract
- 2 Aspiration of respiratory tract frequently
- 3 Oxygen by mask or otherwise as indicated
- 4 Respirator as required for respiratory paralysis
- 5 Intravenous fluids as necessary
- 6 Treat complicated pneumonia with antibiotics if present

Prophylaxis

A Adequate sterilization of all canned foods

B Biting from infected foods before for 5-10 minutes

C Disinfection of tin cans with boiled lids or jars with leaking lids

TULAREMIA (code No 010 107)

Tularemia is an infection produced by the gram-negative organism *Fa. tularensis* which is acquired by contact with infected animals or by bite of insect (deer fly, etc.). Local ulceration and regional nontyphoid pneumonia or septicaemia may be present. Diagnosis may be confirmed by isolation of the organism (difficult) by high agglutination test or by skin test. The incubation period 2 to 7 days.

Treatment in addition to giving symptomatic and supportive measures consists of giving one of the following: (1) tetracycline 0.5 Gm. every 6 hours orally for 5-10 days (2) streptomycin 2.0 Gm. I.M. daily in divided doses every 6 hours for 5-10 days (3) chloramphenicol (Chloromycin®) 0.5 Gm. orally every 6 hours for 5-10 days

PLAGUE (code No 010 106)

Plague caused by *Pasteurella pestis* is characterized by a painful bubo with surrounding edema and severe constitutional symptoms. *P. pestis* (code No 361 106) resembles other enteric pneumonias. The diagnosis is based on isolation of organism from blood, sputum or blood.

Treatment consists of giving dihydrostreptomycin 2.0 Gm. daily I.M. in divided doses or tetracycline 0.5 Gm. every 6 hours and sulfadiazine in full dose (see p. 499). Give symptomatic and supportive measures as needed.

Prophylactic measures consist of giving plague vaccine (2 billion organisms per cc) 0.5 ml. i.c. at interval of 7-10 days. The patient should have gamma globulin as fully diluted.

CHOLERA (code No 010 129)

Cholera is an acute dysenteric disease caused by infection of *Vibrio cholerae*. After a short incubation period of 1 to 4 days a period of profound diarrhea ensues usually accompanied by severe dehydration, electrolyte deprivation and shock. Abrupt onset and prostration are the most important clinical features. The organism may be isolated from stools or vomit. Spread is principally by contaminated water or food.

Treatment.A Specific Measures

- 1 Streptomycin 1.0 Gm every 6 hours I.M.
- or 2 Sodium sulfadiazine 5.0 Gm (75 g) in physiological saline solution I.V. followed by 3.0 Gm (45 gr) I.V. every 8 to 10 hours. Oral sulfadiazine may be substituted when vomiting ceases.
- 3 Sodium bicarbonate in equal or double doses should be given with the sulfadiazine when the patient is able to swallow.

B Supportive and Symptomatic Measures

- 1 Human plasma and physiological saline or Ringer's solution I.V. until shock, dehydration and anuria are alleviated. Large amounts may be required.
- 2 1/8 molar lactate solution may be necessary in severe cases to combat acidosis and to prevent sulfadiazine crystal formation in the kidneys. Solutions containing potassium should be given to eliminate hypokalemia after initial shock and dehydration are relieved.

Prophylaxis

- A Cholera vaccine 0.5 cc initially and then 1.0 cc subcut after a 7 to 10 day interval. Repeat 1.0 cc every 4 to 6 months.
- B Rigorous isolation of all cases and careful disinfection of excreta are important. In endemic areas all water and milk must be boiled and protective screening against flies must be secured.

LEPROSY (code No 010 124)

Leprosy is a chronic disease caused by *Mycobacterium leprae* and characterized by aesthetic granulomatous nodules or macular skin lesions. Enlargement of nerve trunk may be noted by palpation. Multiple atrophy may be seen. The organism may be best demonstrated by biopsy but nasal scrapings may reveal its presence.

Treatment consists of giving Thiophene Sodium N.N.D. (Promizol®) 0.25 to 0.5 Gm every 6 hours orally or Glucophage Sodium N.N.D. (Promin®) 4 cc I.V. daily for 3 weeks followed by an interval of rest for one week (continuous for prolonged periods). Hemolytic anemia should be guarded against by frequent blood counts. May also treat with Isoniazid N.N.D. (INH) 5 mg/Kg body wt per day in 3 or 4 doses.

DISEASES DUE TO SPIROCHETES

(Syphilis discussed on p 435)

RELAPSING FEVER

(Louse borne code No 010 1411)

(Tick borne code No 010 1412)

Relapsing fever is characterized by recurring febrile episode of 3 to 5 days duration following bite by an infected tick or louse. Diagnosis may be confirmed by demonstration of Borrelia recurrentis in the blood on direct examination or by animal inoculation.

Treatment consists of giving (1) penicillin 50 000 units I.M. every 3 hours or penicillin procaine 300 000 units I.M. daily for 10 days or (2) tetracycline 0.5 Gm. every 6 hours orally. Chloramphenicol (Chloromycetin®) or oxytetracycline (Tetracyclin®) may be expected to prove effective. Give supportive and symptomatic measures as needed.

LEPTOSPIROSIS (Weil's Disease) (code No 010 142)

Spirochetes are characterized by severe constitutional symptoms, prothrombin time, and a high fever. A history of contact with rats may be obtained (e.g. sewer workers, war hoistmen). Darkfield examination of the blood or urine revealing the Leptospira interrogans is a high titer of specific agglutination in the diagnosis. Aspt. meningitis, pneumonia, and prothrombin time are also caused by leptospires.

Treatment consists of giving tetracycline drug 0.5 Gm. every 6 hours orally or penicillin 100 000 units every 3 hours I.M. However, the role of antibiotics in treatment is still being debated. Give supportive and symptomatic measures as needed.

RAT BITE FEVER (Sodoku) (code No 010 134)

Rat bite fever is caused by Spirillum minus and is characterized by a recurrent fever at the site of inoculation accompanied by regional dermatitis, fever, and a macular rash. The fever is episodic and recurrent.

Treatment with penicillin 100 000 units every 3 hours I.M. or penicillin procaine 300 000 units I.M. every 12 hours orally is the drug of choice. Give supportive and symptomatic measures as indicated.

YAWS (Fimbria) (code No 010 146)

Yaws is an infectious disease produced by Treponema pertenue and is characterized by granulomatous lesions of the skin, mucous membranes, and bones. Yaws is rarely fatal.

Differential

Yaws is distinguished by the following characteristics. The mother yaw is a small papule which later ulcerates and is 3 to 4 weeks

after exposure Six to 12 weeks later similar secondary lesions appear and last for several months or years Late gummatous lesions may follow Visceral involvement is rare The Wassermann and flocculation tests are positive and the spirochetes may be demonstrated by dark field examination

Cleanliness of lesions is most important in treatment Specific measures consist of giving one of the following (1) penicillin procaine 300 000 units I M daily for 7-10 days (2) tetracycline drugs 0.5 Gm every 6 hours for 10 days or (3) dichlorophenarsine (Cloarsen®) 40 mg I V weekly for 3-6 weeks

INFECTIOUS DISEASES OF UNDETERMINED ETIOLOGY

VINCENT'S ANGINA (Stomatitis) (code No. 810.141)

Vincent's angina is an ulcerative infection of the oropharynx of doubtful etiology Fusiform and spirochete infection and herpes simplex virus have been incriminated

Treatment

A. Specific Measures (Probably against secondary infection)

- 1 Penicillin trochea 5000 units to be sucked constantly
- 2 Penicillin procaine 300 000 units daily I M in severe cases
- or 3 Tetracycline drugs chloramphenicol (Chlormycetin®) may reduce secondary infection (see pp. 507 to 510)

B. General Measures

- 1 Sodium perborate or hydrogen peroxide mouth washes
- 2 Correction of oral hygiene by a dentist

INFECTIOUS MONONUCLEOSIS (code No. 010.1301)

Infectious mononucleosis is an infectious disease probably of viral origin with protean manifestations the commonest of which are sore throat and lymphocytosis Duration is occasionally prolonged for months deaths are very rare

Diagnosis

Clinical symptoms are extremely variable Sore throat fever and malaise are common Lymphadenopathy and splenomegaly usually occur Exudate may be present in the throat Initial leukopenia is generally succeeded by the appearance of a raised number of abnormal appearing lymphocytes Sheep cell agglutinins generally appear at some time during the course of the disease but may be transient Various rashes may be noted Infection period and period of infectivity are not established

Treatment

- ##### A. Specific Measures Many agents have been proposed as affecting the course of the disease To date none have proved effective

B General Measures

- 1 Bed rest until afebrile
- 2 Analgesia with Aspirin or codeine if required
- 3 Hot saline or 30% glucose throat irrigations or gargles 3 or 4 times daily may help
- 4 In severely ill patients symptomatic relief may be afforded by corticotropin (ACTH) or one of the corticosteroids (see p 424)

C Treatment of Complications

- 1 Hepatitis (code No 680 130) myocarditis (code No 430 130 9) or encephalitis (code No 930 130) may occur and are treated symptomatically
- 2 Rupture of the spleen (code No 520 130 5) requires emergency splenectomy. Frequent vigorous palpation of the spleen is unwise

Prophylaxis

None

DISEASES DUE TO PROTOZOA

MALARIA

Malaria is an infectious disease which is caused by one of several species of protozoa of the genus *Plasmodium*. It is ordinarily transmitted only by the bite of a mosquito. There are 3 main species infecting man:

- 1 *P. vivax* (code No 010 1571) Causes vivax or benign tertian malaria
- 2 *P. malariae* (code No 010 1572) Causes quartan malaria
- 3 *P. falciparum* (code No 010 1573) Causes fatal malarial fever or malarial meningitis. Probably the most dangerous because of its tendency to ascend to the brain.

Diagnosis

- A History The disease is characterized by paroxysms of chills followed by fever and sweating which may occur every day (quotidian), every other day (tertian), or every third day (quartan). The presence of unexplained fever in patients in endemic areas presumptively indicates malarial fever. Specific antimalarial therapy should be used in such cases even though organisms cannot be demonstrated. Other wise do not diagnose malarial fever unless the organism has been identified.
- B Laboratory Examination The specific diagnosis is made by finding on examination of *Plasmodium* in thin and thick stained blood smears or bone marrow smears.

Treatment

The routine treatment of malaria therapy represents a rational approach based upon biological concepts to this disease. These concepts distinguish the types of malaria requiring specific drug therapy. The principal antimalarials and a discussion of their use follows.

- A Specific Measures** The following is a description of the available antimalarial drugs, dosages, indications, and toxicity.
1. **Chloroquine** An effective agent against all forms of malaria. *Treatment of choice for all forms of malaria during acute attack.* It will terminate *P. falciparum* infection and prevent relapses of vivax malaria when administered in conjunction with primaquine.
 - a. **Therapeutic dosage schedule**
 - (1) Oral Chloroquine Phosphate U.S.P. (Aralen®) 1.0 Gm (15 gr) as initial dose, 0.5 Gm (7½ gr) in 6 hours and 0.5 Gm (7½ gr) daily for the next 2 days.
 - (2) Emergency treatment: Chloroquine hydrochloride 0.203 Gm (3.5 gr) of base I.M. repeated in 6 hours if necessary. Follow with oral therapy as soon as possible. It is not necessary to administer this drug I.V. since an effective blood level is rapidly attained by the I.M. route.
 - b. **Suppressive dosage** 0.5 Gm (7½ gr) chloroquine di-phosphate weekly, taken on the same day each week.
 - c. **Toxicity** There are few toxic symptoms from this drug when given in the above mentioned doses. These are mild headache, pruritus, anorexia, blurring of vision, malaise, and urticaria. If symptoms become severe, stop drug and give ammonium chloride 4.0 Gm (50 gr) Stat and 1.0 Gm (15 gr) every 4 hours as difficult to promote excretion of the drug.
 2. **Amodiaquine Hydrochloride N.N.D. (Camoquin®)** closely related to chloroquine chemically and pharmacologically.
 - a. **Therapeutic dosage schedule** Give 0.6 Gm (9 gr) of base first day and then 0.4 Gm (6 gr) daily for next 2 days.
 - b. **Suppressive therapy** 0.4 Gm (6 gr) of base once weekly.
 - c. **Toxicity** Mild, similar to chloroquine.
 3. **Quinine** Oldest specific antimalarial drug. Useful in the acute attack of all types of malaria. Prior to the advent of new antimalarial agents, quinine was the drug of choice in the therapy of malaria. If none of the new and more effective agents are available, quinine is still a useful antimalarial drug in a resting or acute attack.
 - a. **Therapeutic dosage schedule**
 - (1) Quinine Sulfate U.S.P. B.P. 0.6 Gm (10 gr) tid orally for 5 to 7 days.
 - (2) Quinine Dihydrochloride N.F. B.P. 0.65 Gm (10 gr) in normal saline glucose salt mixture or plasma injected I.V. VERY SLOWLY (not more than 50 mg of salt per minute) at 6 hours intervals. Give no more than 3 injections in 24 hours. May also be administered by I.V. drip at the rate of 2.0 Gm (30 gr) in 24 hours. Follow with oral therapy as soon as possible.
 - b. **Suppressive dosage** Quinine sulfate 0.306 Gm (5.10 gr) daily while in endemic area.
 - c. **Toxicity** Quinine in the above dosage may cause cinchonism (tinnitus, vertigo, deafness, headache, visual disturbance) in some individuals. (The drug is a

far less effective than the putrid and suppressive agents than some of the newer and less toxic preparations. The possibility of bloodwater fever arising directly or at the cessation of therapy appears to be higher in quinine treated cases.

- 4 Proguanil hydrochloride (Paludrine®) Although not a definitive agent for the treatment of the acute clinical attack this is a good suppressive drug for all forms of malaria. It has a tendency to produce resistance.
 - a Dose 1 to 2 g daily for prophylaxis; 0.1 Gm (1½ g) daily for prophylaxis; 0.3 Gm (5 gr) once weekly
 - b Toxicity Slight. Large doses cause nausea, vomiting, diarrhoea and mild haematuria.
 - 5 Pyrimethamine (Dapron®) although not recommended for the treatment of the clinical malaria is a effective agent for prophylaxis.
 - a Dose 1 to 2 g daily for prophylaxis; 25 mg (38 g) weekly taken on the same day of each week. For children give 12.5 mg (3/16 gr) weekly (may be dissolved in syrup).
 - b Toxicity Very low at recommended dose.
 - 6 Primaquine Phosphate U.S.P. Research work has shown this drug to be the most effective against the merozoites of the Plasmodium vivax. This drug is employed to eradicate the disease rather than to treat the clinical attack. It will prevent relapse in the majority of cases.
 - a Dose 1 to 2 g daily for prophylaxis; 25.4 mg (15 mg base) daily in divided doses for 14 days followed by at least 2 weeks with chloroquine phosphate or amodiaquine if indicated.
 - b Toxicity Chloroquine resistance is not a problem. Severe hemolytic reactions can occasionally occur particularly in Negroes. Watch for fall of haemoglobin concentration during treatment.
- B G al M res. The onset of the first attack is not different from that of a typical malarial fever. The symptoms are usually mild and in the immediate period of infection.

AMEBIASIS (cod N 010 151)

Amoebiasis is a chronic infection caused by the protozoan Entamoeba histolytica. Although not a primary infection, it is often a life-threatening disease to the individual. The term amoebiasis applies to both the systemic and the local disease. The term amoebiasis is used to describe the systemic disease. The term amoebiasis is used to describe the systemic disease. The term amoebiasis is used to describe the systemic disease.

A Am bi Coliti (Dysentery) (cod No 660 151)

- 1 Acute With diarrhoea and other gastrointestinal symptoms
- 2 Chronic Without diarrhoea

B H p ti Am bi al

- 1 Hypertensive Amoebiasis (cod No 680 151)

- 2 Liver abscess Acute or chronic (code No 880 151 2)
- C Amebiasis of other organs (uncommon) lungs brain etc may be involved
- D Asymptomatic amebiasis (carrier state?) may actually be a latent phase of the disease

Diagnosis

A history of exposure to infection in endemic areas or in epidemics is important but remember that there is an almost world wide incidence of the disease. A past history of dysentery or diarrhea (especially known amebiasis) is exceedingly valuable

A Amebic Colitis

- 1 Bloody diarrhea is frequently present during acute phase
- 2 Stool examination
 - a Blood and mucus may be present but there is usually comparatively little pus (cf bacillary dysentery p 278)
 - b Cysts and/or trophozoites of *E. histolytica* should be demonstrated. Repeated stool examinations of carefully collected fresh specimens are necessary for this
- 3 Fever and leukocytosis may be present
- 4 Multiple vague gastrointestinal symptoms may be present in the chronic phase of amebic colitis
- 5 Sigmoidoscopic examination may reveal amebic ulcerations

B Amebic Hepatitis and Liver Abscess

- 1 Symptoms of dyspepsia may be present
- 2 Signs and symptoms of adjacent right chest involvement may be present
- 3 Liver may be enlarged and is frequently tender and even painful. Occasionally a localized mass is palpable when abscess is present. Liver function tests may be abnormal
- 4 Mild to moderate leukocytosis and fever may be found
- 5 Stools do not usually show *E. histolytica*
- 6 X rays may reveal alteration of contour of diaphragm, hepatomegaly or right lower chest involvement
- 7 Material (anchovy paste like) may at times be aspirated from a fully localized abscess mass

C Amebiasis of Other Organ Diagnosis is difficult and is possible only by maintaining a high index of suspicion of specific organ involvement (based on clinical manifestations) in patients with known or suspected amebiasis

D Asymptomatic Amebiasis This diagnosis must be reserved for cases in which routine stool examination reveals cysts of *E. histolytica* but clinical findings (including sigmoidoscopic examination) are completely negative

E Therapeutic Test A therapeutic trial of anti amebic drugs particularly chloroquine or emetine may be warranted

- 1 If diagnosis is doubtful after case of investigation and amebiasis (especially hepatic) is clinically suspected
- 2 If fulminant diarrhea is present and diagnosis is clinically suspected but cannot be established and no other organism can be found

Treatment

A General Measures

- 1 Report case to local health authorities
- 2 Bed rest is required for certain patients

With Frank

dys tery hepatic o oth r non enteric involve me t and all patients receiv g emetine therapy (See below)

3 D t

a If diar hea is p esent follow the dietary m asu es as outlin d fo n nspc ific di rhea (see p 258)

b If ther is hepatic da ease follow the dietary measures with ed d r th management of chr ni hepat c dis ease (see p 281)

c Iron salis ho ld b given if a emia i pres nt (s e p 219)

B Ac te or Ch nic Amebic Dys tery

- 1 Sp cif d g In the p ence of dys nt ry it is p obably safe to assum that organisma have invaded extra intestin l tiss es With this n mind n ad qu te co rse of th rapy should in lud not only an gent eff ctive against intestinal form but also a d ug which is effect v in th extra int ti al tis es (ee table b low)

Eff ctiveness of Anti amebic Drugs

	Chlo o quin	Em tine	Ca ba on Milbis®	Vio form®	E yth o mycin	Fum gillin
I t st l						
O gan m	±	±	+	+	+	+
Ext sine tinal						
O ganl m	+	+			±	±

Combi at ons of hlo quine (o emetin) and an rs nical (a ba s n or Mil bis®) or an iodine cont ining mp und (Vioform®) ar ommonly mploy d If th organisma prov e istant ythromycin or fum gillin an b tri d D ag s e giv n below

Chl roquine Phosph t U S P (Ar len®) 0.5 Gm (7½ gr) (o 0.3 Gm of th b s) b i d f 2 day follow d by 0.5 Gm (7½ gr) d ily f 7 to 10 d y

- b Em tin Hyd hl rid Inj ct U b P B P 65 mg (1 gr) da ly sub t f r 6 d ys will cont ol a ute symp t m deradic tes infe ti n in 15 p cent of se Em tin b b en r plac d by less toxic and qually ef f tive g nt su h s hlo quine (see ab v)

C bar one U S P B P 0.25 Gm (3¾ gr) i d ily f r 7 to 10 day o Gly obs 1 N F (Milb ®) 0.5 Gm (7½ gr) i d fo 7 day

- d Iodo chlorhyd o yq in U S P (Vi f rm®) 0.2 Gm (¾ g) t d o ily fo 14 d ys E yth my U S P 15 mg p Kg body w ght d ly fo 10 to 14 d y This ant bot i ffectiv fo th t m t f a team blas

f F m gli N N D (F m d l®) 0.5 to 1 mg p Kg body w ght d ly f 10 t 14 days Empl y d f r d ug ef ti a s of hro cam bi s

- 2 Ev luatio of th py Following ompl tion of th py w th w kade m e tool on th e s c s ive d ys a If till pos t v epe t bow o r of t atm nt d r exami fo f at ou se b If n gative giv no f th r treatm nt R e m s stool

at 4 week intervals until a total of 12 specimens have been found to be negative

3 Toxic reactions of the anti amebic drugs

- a Emetine hydrochloride *Contraindicated in myocardial disease* Nausea vomiting muscular weakness neuritis myocarditis and prostration may occur Special observations and precautions include the following
 - (1) Bed rest without lavatory privileges
 - (2) Blood pressure determination b i d
 - (3) Pulse determination q i d
 - (4) Daily examination of patient
 - (5) Ecg before and after course of therapy
 - (6) Withd aw drug in the event of toxicity
- b Arsenic containing compounds (Carbarsone Milibis®) *Contraindicated in hepatic disease* Nausea vomiting colic diarrhea and dermatitis may occur Daily inspection for toxic symptoms is necessary Stop drug in event of toxic reaction
- Iodine containing compounds Iodochole hydroxyquin (Vioform®) *Contraindicated in renal disease* Gastrointestinal upsets and diarrhea may occur Daily inspection for toxic effects (uncommon) is necessary Stop drug in event of reaction and resume after few days with smaller doses

C Hepatic Amebiasis

1 Measures for hepatitis

- a Chloroquine Phosphate U S P (Aralen®) is the drug of choice in hepatic amebiasis since it has proved to be effective in emetine resistant cases and is much less toxic Like emetine this drug has rather feeble intestinal effects it is therefore necessary to follow the course of chloroquine with Vioform® carbarsone or one of the antibiotics notably erythromycin or fumagillin (see above)
- Dose Chloroquine diphosphate 0.5 Gm (7½ gr) (or 0.3 Gm of the base) b i d for 2 days followed by 0.5 Gm (7½ gr) daily for 12 to 18 days
- or b Emetine hydrochloride injection 60 mg (1 gr) subcut daily for 8 day If chloroquine is not available or fails to provide a satisfactory therapeutic effect
- c Erythromycin and fumagillin now under trial may prove to be equally effective against both hepatic and intestinal amebiasis
- d General supportive measures should be instituted as for infectious hepatitis (see p 279) A 2 week rest period may be followed by a period of treatment After the patient has convalesced from his hepatic disease further anti amebic drug therapy might be considered in an effort to eradicate the intestinal infection

2 Measures for liver abscess

- a Treat as for hepatitis (see above) If patient responds to chloroquine or emetine treatment follow up with other amebicides for a period of 2 weeks (see p 481) A repeat course alternating these drugs may be necessary after a rest period of one week
- b Small abscesses If patient responds is treated with use combined and then alternately

o met e and othe drug therapy (see abo)

c La g ab e s If pat ent d es not show defi ite spon e to metin o chloroquine t tme t

(1) Co t e t eatment nd r fully att mpt to localiz ab ce s s te by physical and ray f nding Aspirate under s pt c o d tions (p efe bly op ating room) and repeat a piration if n c ssa y Avoid open dr i age nless absce s is econd rily infe ted

(2) Co t e d ug the py (ee above) until evid n e of both h pat a d intestinal d s se i dicat d

d S condarily inf ted absce s

(1) If asp rat d mate lr veals pus and o ganisms (by sm ar and cultu) it m y b necs ary to stabl h pend d inge (by extra ro s app oaches)

() Chemotherap uti ag nts sho ld be employed in the e a es (s pp 496 514)

(3) Compl te u se of nt am bic the apy s for hepatit o liv r ab es if ndic ted (ee above)

3 Ob erve fo tox r ct ns (s ep 482)

D Am bi f Oth ORG s Sp if c the py as fo ut r h nic me b c dys te y

E A ympt m tic Ameb asis (C r r State)

1 F ll o s Som lnt a f l that ameb sis lway h s systemic as w ll s l cal eff ct a d r omme d a full cou s of th py fo a ut o hron c dys te y (p 481)

2 Simple o al mbulatory t e tme t co id ed s tisa tory by th r lni ans

a Ca ba sone 0 5 Gm ($3\frac{3}{4}$ g) t i d o lly fo 7 d y

b V of m[®] 0 25 Gm ($3\frac{3}{4}$ g) t i d o lly f 14 d y

c F ll w p sto l xamination sho ld b perf med as fo a t e am b c nf t (s ep 481)

F Follow p Pat e ts sh ld b f ll wed by lni al s d l bo t ye amin to all t on oc a m h month fo 3 to 6 months following th apy Exami ation sho ld in l d s g m ides py d study ff h stool on 3 s sse ve d ys (p ef ably at lea t l following aline athar) R pe t ex aminati sho ld be p form d w thin a ya if nec sary Ne d f h moth rapy ma t b ba d pon a tual d mont a tio of am b the th n me p se ce of ymptom (e g h ond d h a) Con de th po sibi lty f mpli ations f th d a i e s onda y inf t i t tion of bow l f om ch moth py p y hic traum et wh n pe sistent sympt m a t b t t i d by labo t ry f ding Avoid ov t e t m t w th m ti o oth dr g

GIARDIASIS (cod No 604 155)

Gard is m if st d by e nt pl o d s of w tery s m i d o b lky and ft f l m lling tool Am ld t h l h l y t t is may o r The p f c d agr i is m de by d m r tion f G dia lambli (t i tatin li) n th tool

T atm t con i t of gi ling Quina i Hydro hl id L S P (At b e[®]) 0 i Gm ($1\frac{1}{2}$ g) t d fo 5 day a d Amodaq i Hyd hl id N N D (C moq [®]) ingl d e f 0 6 Gm (9 g)

Repeat if necessary Repeat stool examination after one week to determine efficacy of treatment

TOXOPLASMOSIS (code No 010 1577)

Toxoplasmosis is a disease of man and animals caused by *Toxoplasma gondii*. It is most frequently encountered in the newborn who acquire their infection in utero. Active toxoplasmosis is rare in adults although inapparent infections recognized by serologic tests are not uncommon in the general population. Infants and young children with the disease show evidence of chorioretinitis, cerebral calcification, hydrocephaly or microcephaly and profound psychomotor disturbances. Convulsions may occur. Little is known of the mechanism of transmission of the parasite from one host to another. Clinical diagnosis is most frequently established on the basis of the presence of cerebral calcification and chorioretinitis. In fact, the latter may be the most important single manifestation of acquired toxoplasmosis. For laboratory confirmation the organisms may be demonstrated in smears of blood, bone marrow, C.S.F. or exudate from serous cavities. *Toxoplasma* can usually be demonstrated in laboratory mice following intracerebral and intraperitoneal inoculation of fluids, tissues or exudates. The complement fixation test and the microchemical dye test of Sabin and Feldman are the most useful diagnostic procedures.

There is no effective treatment although combined therapy with sulfadiazine and the antimalarial pyrimethamine (Daraprim®) has shown encouraging results in murine toxoplasmosis and in a limited number of human infections.

DISEASES DUE TO METAZOA

TRICHINOSIS (code No 255)

Trichinosis is caused by the ingestion of raw or improperly cooked infected pork. It can also be traced to other individuals who have consumed the same infected food. The incubation period is from 3 days to 4 weeks.

Acute manifestations may be very mild or may be fatal. Nausea, vomiting, cramps, diarrhea, flatulence and constipation occur early and are followed after a few days by fever, chills, weakness, rash, periorbital and dependent edema, splinter hemorrhage, pain and tenderness in muscles and eosinophilia. A cyclical or peripheral neurological involvement may be present. Headache is a prominent feature. A delayed reaction to the trichinella skin test (noted only after 12 to 24 hours) occurs early in the disease (3 days to 7th day).

Chronic infection is manifested by vague weakness and other symptoms referable to multiple organ systems. Eosinophilia is often marked. Muscle biopsy may demonstrate organisms. An immediate reaction to the trichinella skin test (noted after 5 minutes) occurs late in the disease (from 17th day on).

Treatment is supportive and symptomatic. Severe acute cases require hospitalization and excellent nursing care. Cortisone

(ACTH) and the corticosteroids provide effective relief for the acute symptoms. A reduction of the eosinophil count and disappearance of fever and splinter hemorrhages if present and a general improvement in the clinical status of the patient are guides which should be employed to determine the efficacy of treatment. In the acute stage treat with relatively large doses of either drug for the first 24 to 48 hours (see p 423). In the subacute stage therapy may have to be continued for several days or weeks; prevent recurrence. Dosage is reduced keeping symptoms under control (see p 423).

ASCARIASIS OR ROUNDWORM DISEASE (code No 650 241)

Infection with *Ascaris lumbricoides* may cause no symptoms or only mild fluid gastrointestinal and nervous symptom. Occasionally general edematous reaction may develop rarely a carispermia may result. The specific diagnosis is made by finding the worm's eggs or dead worms in stools or vomitus or by observation of the adult worm passing from nose or mouth.

Treatment.

A H xyl or in l U S P (d g of ch i f ad lts)

- 1 Intestine purgative. Give 30 Gm (1 oz) magnesium sulfate in water or 240 cc (8 oz) of solution of magnesium citrate at the night before drug therapy.
- 2 A light meal is given the preceding evening and then no food until at least 5 hours after taking the hexyl esor in l.
- 3 Alcohol is contraindicated before and during treatment.
- 4 Hexyl or inol 5 hard gelatin capsules 0.2 Gm (3 gr) (capsules) (total 1.0 Gm 15 gr) is given in the morning on an empty stomach. These are to be swallowed whole not chewed. Do not feed to children under 6 years of age 0.4 Gm (6 gr) 6 to 8 years 0.6 Gm (9 gr) 8 to 12 years 0.8 Gm (12 gr).
- 5 Purgation. Two hours later give 30 Gm (1 oz) magnesium sulfate in water to remove the worms from the bowel. Repeat 2 hours later if necessary for purgation.
- 6 Stool examination should be made once weekly for on 3 consecutive days to determine efficiency of treatment.
- 7 Treatment may be repeated in 3 days if necessary.

- B P p zi Cit te, N N D (Ant par®) (D g of cho f child n) Each one of the syrup contains 100 mg p p z in h anhydrous tablet equivalent: 250 and 500 mg (give 5 to 7 days or as follows (daily dosage))
- | | |
|-------------|------------------------------------|
| Up to 15 lb | 1/2 tsp or one 250 mg tablet daily |
| 15 to 30 lb | 1/2 tsp or one 250 mg tablet b i d |
| 30 to 60 lb | 1 tsp or one 500 mg tablet b i d |
| Over 60 lb | 2 tsp or two 500 mg tablets b i d |

- C Di thyl arb m in Cit t U S P (H t zan®) Give 3 to 6 mg /Kg body weight orally 3 times daily for 7 to 11 days. A syrup preparation containing H t a n® powder in a concentration of 30 mg / cc is recommended for small children. Administer 12 mg /Kg body weight once a day for 4 days or 8 to 10 mg /Kg body weight twice daily for 7 to 10 days. When H t a n® is used for radiolysis of *Ascaris lumbricoides* peritremis following a posttreatment purgation is not necessary.

- D Oil of Chenopodium and Tetrachloroethyl n May be used if other preparations are ineffective or not available. Caution: Tetrachloroethylene stimulates activity of ascaris and may result in bowel obstruction. A preliminary course of hexylresorcinol is advised before using the combined method.
- 1 Follow procedure of treatment as mentioned above for hexylresorcinol.
 - 2 Oil of chenopodium 0.3 cc ($4\frac{1}{2}$ w) capsule and tetrachloroethylene three 1.0 cc (15 w) soluble gelatin capsules (total dose 3.0 cc 45 w) are given together and followed by purgation as mentioned above.

ANCYLOSTOMIASIS OR HOOKWORM DISEASE (code No. 650 243)

Most commonly caused by *Necator americanus* or *Ancylostoma duodenale*, the disease occurs when a sufficient number of the worms are present in the intestine. It is manifested by fatigue, weakness, dyspnea, palpitation, anorexia, perverted appetite, weight loss, and a mild microcytic anemia. Ground itch, an erythematous or maculopapular or vesicular pruritic dermatitis, may develop at the site of penetration of the skin by the larvae. Specific diagnosis is made by finding the eggs in the stool.

Recent work indicates that symptoms are most often due to a coexisting deficiency disease. Correction of the malnutrition by adequate dietary means and of the anemia by the addition of iron appears to alleviate or remove the manifestations in the absence of specific treatment for the hookworm infection.

Treatment.

- A General Treatment. Estimation of the need for treatment should be based upon quantitative counts of the eggs in the stools. There is no indication for treating light infections, particularly after completion of previous therapy. It is often impossible to completely eradicate the infection.
- 1 Correct malnutrition. Provide an adequate protein diet with supplementary iron medication (see p. 218).
 - 2 Rule out possibility of coincidental ascariasis. If ascariasis is present or when diagnosis falls within a delimited give preliminary hexylresorcinol as prescribed for ascariasis (see p. 485). Tetrachloroethylene stimulates ascaris activity which occasionally results in intestinal obstruction. If large numbers of hookworms are still present following the administration of hexylresorcinol wait one week following the last dose and give tetrachloroethylene.
- B Specific Treatment.
- 1 Tetrachloroethylene U.S.P. B.P. (drug of choice). First correct malnutrition and anemia. Tetrachloroethylene is contraindicated in patients with alcoholism, chronic gastrointestinal disorders, severe constipation, hepatic disease, and in patients undergoing hypomyelination. Correct these conditions before giving treatment.
 - a Initial purgation. Give 30 Gm (1 oz) magnesium sulfate in water or 240 cc (8 oz) magnesium citrate solution the night before drug therapy.

- b The meal of the preceding evening should be light no food should then be taken until the patient gets on after treatment has caused a copious bowel movement. Alcohol and fatty foods should not be taken for at least 48 hours prior to drug therapy.

Tetrahydrocortisol 10 cc (15 mg) soluble gelatin capsules (total 30 cc 45 mg) should be given in the morning on an empty stomach.

- d Purgation 2 to 3 hours later give 30 Gm (1 oz) magnesium sulfate in water to remove worms from the bowel. If no purgation results within 4 hours repeat 30 Gm (1 oz) magnesium sulfate enema or any means to hasten evacuation.

Examine stools a week later on 3 successive days to determine efficacy of treatment.

- f Repeat treatment 2 weeks later if indicated.

- g Ferrous sulfate 0.2 to 0.3 Gm (3.5 gr) tid po may be given if anemia.

- 2 Hydroxyresorcinol. May be used if tetrahydrocortisol is contraindicated ineffective or not available (see p 485).

STRONGYLOIDIASIS (code No 604 256)

Infection with *Strongyloides stercoralis* manifested by gastrointestinal large bowel disturbances watery mucus stools and eosinophilia. Fever, weakness and weight loss may be present and strongyloid pneumonia may occasionally occur. Specific diagnosis made by demonstration of motile threadiform larva in the stool (coproctum).

Treatment consists of giving gentian violet 60 mg (1 gr) each twice a day until 3.3 Gm (50 gr) have been taken. In 1½ hours not eat or drink. Old children should receive adult dose. Young children a daily dose of 10 mg (¼ gr) if achy of proppant (not chronological) age not exceeding adult dose. For refractory cases 25 of a 1 per cent aqueous

solution of gentian violet medicinal powder may be instilled into the rectum daily. Red dosage by ½ rectally temporarily discontinue treatment if any of the following appears: severe proppant discomfort, flatulence, severe nausea and/or vomiting or violet discharge from the rectum.

ENTEROBIASIS OR OXYURIASIS (code No 604 242) (Pinworm or Seatworm)

Infection with pinworms usually presents itself and often severe perianal pruritus usually more intense at night and frequently in the itching sexual member of a household or institution. It occurs most commonly in children. There are often a variable vague gastrointestinal and nervous symptoms. The specific diagnosis is made by demonstration of the eggs of *Enterobius* (Oxyuris) vermicularis in the perianal or perineal skin (Swathill's cellulose tape method or by hand). Eggs are not commonly found in the stool.

TreatmentA General Measures

- 1 Examine for and treat all infected members of the family and other groups of close contacts since infection from non treated contacts is frequent
- 2 Instruct patient and/or household members with respect to
 - a Careful washing of hands with soap and water after defecation and again before meals
 - b Keeping fingernails trimmed close and clean
 - c Voluntary abstinence from scratching of involved areas
 - d Apply carbolated petrolatum to anal region following every defecation Thoroughly wash anal region in morning with soap and water
 - e Daily morning showers or stand up bath with acapy water
 - f Scrubbing of toilet seats with soap and water daily
 - g Cleaned boiled bed linens 2 times weekly
 - h Use of pajamas (or sleepers for children) to prevent manual contact with anal region during night
 - i Discourage patients from putting hands into mouth
 - j Raise temperature of sleeping rooms as high as possible for one hour daily then a rather roughly

B Specific Measures (listed in order of effectiveness)

- 1 Piperazine Citrate N N D (Antepar®) (syrup or tablets)
Each cc of the syrup contains 100 mg piperazine hydrochloride tablets are equivalent to 250 or 500 mg Administer daily for 7 days follow with a rest period of 7 days and then administer again for another 7 days Use dosages given for Ascaris p 485
- 2 Oxytetracycline (Terramycin®) Oral drop oral suspension or capsules Give 10 mg per lb body weight daily divided into 3 doses administer daily for 7 days
- 3 Methylthiouracil Chloride U S P Crystal Violet B P (4 hour enteric coated tablet) 1 mg per lb body weight divided into 3 daily doses Administer daily for 8 days follow with a rest period of 8 days and then give again for 8 days

TAPEWORM INFECTIONS (code No 604 261)

(Pork Beef Fish Dwarf or more
rarely Dog or Rat Tapeworms)

Tapeworm infections are caused by consumption of contaminated raw or incompletely cooked pork beef fish water fowl or other contaminated food The acute phase (usually after a long incubation period of 3 to 4 months) is manifested by diarrhoea fever leukocytosis and eosinophilia The chronic phase is manifested by vague gastrointestinal and CNS symptoms mild to severe anaemia and gravid proglottids in feces or underclothing Specific diagnosis of infection with Taenia solium (beef) Taenia saginata (pork) and Dipylidium caninum (dog) is made upon appearance of gravid proglottids in feces (occasionally by eggs in feces) Specific diagnosis of infection with Diphylobothrium latum (fish) Hymenolepis diminuta (rat) is made upon appearance of eggs in feces

T e a t m e n tA S p e c i f i c M e t h o d

1 A p p o i n t m e n t O l e o e s s e n t i a l U S P E t r a c t o f M a l e F e n B P
C t r a i n d i c a t i o n s S e v e r e c a r d i a c h e p t i c o r e a l
d a s c o t i p a n a u t e o c h i c g a s t r o e t e r t a
f e b r i l e a t t e s p r g n a y a n d i n f a y

b L w r e s d u e f t f r d i e t f 24 t o 48 h o u s p o t o d u g
t h e a p y A l c o h o l i s o t r i n d i c a t e d

c M g n u m s l i f a t o r o d i m u l f a t e 15 t o 30 G m (1/2
t o 1 o z) i n w t e r i g n t h e n i g h t b e f o r e t r e a t m e n t
O m i t b r e k f a s t o n m o r n i n g o f t r a t m e n t

d A d m i n i s t e r o l e o i n o f a s p d u m i g e l a t i n c a p u l i n
t h r e q u i d e s a t h a l f h o r i n t e v l s a c h d e o n
t a i n g f m 0.5 t o 1.2 G m d e p e n d g o n w e i g h t o f t h
p a t i e n t C h i l d r s h o u l d e l v i m i m p e y e a f a g e
*The drug should be fresh and not dispensed from
bottles which have been opened for some time*

e M a g n e s i u m u l f a t e o r o d i m u l f a t e 15 t o 30 G m (1/2
t o 1 o z) i n w t e r g e n g n t w o h o u s f o l l o w i n g
a d m i n i s t r a t i o n o f l a s t c a p s u l e N o f o o d s h o u l d b e p e
m i t t e d t i l t h e b w e l m o e c p o u l y

f R e p e t e o u s o f t a t m e n t n o t l e s t h a n 7 d a y s i f
s a r y

g A l t e r n a t e m e t h o d o f d m i t a t o f o l e o n o f a s p d
i m g

A p p o i n t m e n t i n 4

M o d i f y f a l a 30 cc

C o n t a i n e d b u t i n
o f a s p d u m u l f a t e 30 cc

G i t h m u l a o l l y b y d o d l i b e i n o n e d
m i n i s t r a t o P o s t t r a t m e n t p g t i t n s y
O n h a l f t h d o s a g e i f t y f o c h i l d r e f h o o l
g

h T h p a t i e n t s h o u l d h a v e h a t i o n b e f
w m w t r o f i l t a t e p c m e n t a n d d e t a t t n
o f t h h e d a n d p r o g l o t t d T o l i p p h o l d b d s
p o d o f e p t l y E m m a l l s t o o l s p s d d i g
t h e e x t 24 h o u r i n o r d e r t o o v t h e w r m h d f
p f f c o m p l t d i c a t

2 Q u a n t i t y o f H y d r o c h l o r i d e U S P M p c H y d o
h l o d B P (A t a b n o)

a O n d a y p e d i g t e a t m e n t p a t i e n t s l i g h t l o w
r i d e i n h a n d s p p e

b S o d i u m b i c a r b o n a t e m a g n e s i u m b i c a r b o n a t e 15 t o 30 G m (1/2
t o 1 o z) a t t a t i t h e n i g h t b e f o r e t r a t m e n t
O n m o r n i n g f t a t m e n t o m i t b e k f t G v e p h o
b a r b i t 30 t o 80 m g (1/2 t o 1 1/2 g) d e p e n d i g u p n
t h e w e i g h t a n d a g o f t h p a t i e n t O h o l t a d m i n
t a s a n g l e d e 0.5 t o 1.0 G m (1/2 t o 1.5 g)
(d e p e n d g o w e i g h t o f p a t i e n t) o f q u i n a c r i n e h y d o c h l o
r d l o n g w t h a n e q u a l a m o u n t o f s o d i u m b i c a r b o n a t e t o
c o n t r a i n a u c a a n d v o m i t i g

A m o r e e f f e c t i v e r e m o d a l o f t p e w o m t h a n i s p o s s i b l e
w i t h o l m d a l i i t a b l i t f o r m m y b e a c c o m p l i s h e d
b y d m i n i s t r i g q u i n a c i n h y d o c h l o i d t h r o g h a d u o
d l i t u b e

- d Sodium sulfate or magnesium sulfate 15 to 30 Gm ($\frac{1}{2}$ to 1 oz) in water is given 2 hours following therapy to rid the intestine of the parasite. No food should be permitted until the bowel moves copiously.
 - e Repeat course of treatment after 7 days if necessary.
- 3 Hexylresorcinol U.S.P.
- a 1.0 Gm (15 gr) in 20 cc water introduced into the duodenum by a tube. Follow in 2 hours with a sodium or magnesium sulfate purge. Examine stools for head of worm.
 - b Crystoids anthelmintic as administered in ascariasis (see p. 485) is the drug of choice for the treatment of light infections with *Hymenolepis nana* (dwarf tapeworm). For heavy infections use quinacrine hydrochloride as for treatment of *Taenia saginata* infections.

B General Measures Hospitalization is recommended for the treatment of persons with tapeworm infection. Successful removal of the parasite can only be accomplished if there is cooperation on the part of the patient, the clinician, and the laboratory personnel. Proper pretreatment preparation of the patient and adequate postpurgative examination of stools for the head of the tapeworm is necessary.

SYSTEMIC MYCOSES

Mycotic infections are caused by a variety of fungi and have wide geographic distribution. Although their incidence is rather low in certain parts of the world some of them occur quite commonly. *Coccidioidomycosis* in the San Joaquin Valley of California. Their clinical manifestations are exceedingly variable with some resemblances to the granulomatous diseases.

COCCIDIOIDOMYCOSIS (code No. 012 219) (Pulmonary code No. 360 219)

Coccidioidomycosis or Valley Fever is an infection due to *Coccidioides immitis* which is found in the Southwest United States, Mexico, and Central and South America with sporadic cases in Italy and Hawaii.

Diagnosis

- A Primary Form** Involves the bronchi and lung and may manifest itself by fever, cough, erythema nodosum, and pleurisy with effusion. X-ray of the lungs during the primary disease shows patchy soft infiltration which clears. Subtle nodular shadows may persist. Thin-walled cavities with little or no surrounding infiltration may develop and remain for months. The sedimentation rate is elevated. Organisms may be found in the sputum by culture and the skin test may be positive after 10 to 14 days. Complement fixation and precipitin test are helpful in diagnosis and may aid in determining the prognosis of infection.
- B Chronic or Crateriform Form** 0.2 percent of all primary cases progress to the granulomatous stage involving the lung, chest wall, or other structures. The finding of the organisms in infected tissue or in the discharge from the lesion in this stage makes the diagnosis. Prognosis in this form is poor.

Treatment

- N specific therapy is known for either form of the disease
- A Primary Form Bed rest and symptomatic care until process has subsided
- B Chronic Form Treatment entirely symptomatic Potassium iodide is of no value and may even be dangerous

ACTINOMYCOSIS (Regional code No 0 202)

NOCARDIOSIS (Pulmonary code No 360 201)

Actinomycosis is world wide in distribution and caused by a aerobic actinomyces Actinomyces israelii Nocardia is caused by variety of aerobic type belonging to the genus Nocardia (e.g. Nocardia asteroides)

Diagnosis

- A Actinomycosis The principal lesions are multiple abscesses sinuses and fistulas through which discharge a grainy pus like material containing sulfur granules A large portion of the body may be infected but the head and neck are most frequently involved in which case very little systemic reaction occurs The abdominal viscera may be involved by way of the intestinal tract or the lungs pleura and chest by way of the respiratory tract In the latter two forms the disease may be symptomatic of the systemic infection accompanied by high fever The finding of the typical sulfur granules in the lesion is diagnostic
- B Nocardiosis may resemble classical actinomycosis with the production of characteristic granules or produce a peripheral pulmonary inflammation of the lungs complicated with extension to brain and meninges Various species of Nocardia cause infection of the subcutaneous tissue with characteristic involvement (mycetoma)

Treatment

- A Actinomycosis The treatment of actinomycosis must frequently be continued for weeks or months
- 1 Penicillin is the drug of choice An initial dose of 120 000 units should be followed by 80 000 units every 4 hours The dose of much higher doses is essential and by some clinicians daily intramuscular injections of 600 000 units for 6 weeks followed by a further 2 weeks of oral penicillin
 - 2 Sulfadiazine The concentration of sulfadiazine has been found to be fatal in certain instances
 - 3 Potassium iodide With some patients the above combination should be supplemented by iodide therapy but treatment must be continued in the severe patient Give potassium iodide 5 g 5 times a day U.S.P. 5 drops 4 times a day gradually increased until reactions occur at a rate of 20 drops a day is recommended This may be further increased and as much as 100 drops a day may be used if the patient still reacts then do so If reactions occur stop drug and begin again at a low dose gradually patient if may be necessary to start with 1 or

2 drops daily and gradually increase dose

- B Nocardiosis Treat as for actinomycosis. The disease producing aerobic *Nocardia* are more susceptible to the sulfonamides than is *A. Israeli*. Treat with sulfadiazine alone or combine with sulfamerazine. Maintain blood levels of 10 to 20 mg per 100 cc until infection is controlled. Patient should be maintained on sulfo amide therapy 3 to 4 months following apparent cure.

BLASTOMYCOSIS

(Generalized code No 012 200) (Of Skin code No 110 217)

North American blastomycosis chiefly noted in North Carolina, Illinois and Mississippi is due to *Blastomyces dermatitidis*. The South American variety is due to *B. brasiliensis* and is found mainly in Brazil.

Diagnosis

The North American variety produces granulomatous ulcerating lesions of the skin and tuberculosis like lesions of the lungs and at times of other structures. The cutaneous form usually is without systemic symptoms but the pulmonary form usually has symptoms not unlike those of pulmonary tuberculosis.

The South American variety involves the mucous membranes of the mouth, the skin, the lymphatic tissue and the viscera.

The diagnosis of either variety depends upon microscopic recognition of the characteristic yeast like fungus in tissues or exudates followed by isolation of the organism in culture.

Treatment

- A Potassium iodide therapy should be given as for actinomycosis (see above). Before administration of the drug the patient should be tested for sensitivity to the fungus and to the drug. If these cautions are not observed spreading of the infectious process may occur. At the beginning of the treatment only small amounts of potassium iodide should be used. Further more to prevent dissemination of the infection desensitization or immunization with *Blastomyces* vaccine may be necessary.
- B Stilbamidine has proved to be quite effective in the treatment of cutaneous and systemic blastomycosis. Dose: 10 to 200 mg daily not exceeding 2 to 3 mg /Kg daily. It is given slowly I.V. in 5% glucose new tetro saline. A course of 30 days may be required. This drug has to be employed with caution since it is toxic and frequently produces a neuropathy especially of the fifth nerve. A related drug 2 hydroxystilbamidine has been used successfully on patients infected with *B. dermatitidis*. It fortunately does not produce peripheral neuropathy.
- C X ray therapy may be used as an adjunct in the cutaneous cases. The systemic forms are rather resistant to treatment and progress in spite of therapy.

HISTOPLASMOSIS (code No 010 218)

Histoplasmosis is caused by *Histoplasma capsulatum* a small yeast like organism in tissue and a mold like fungus in culture. It has a world wide distribution. It primarily involves the reticuloendothelial system causing enlargement of the liver spleen and lymph nodes and systemic manifestations of fever anemia and leukopenia. However other systems may be involved. Patient from endemic areas often have pulmonary calcification negative tuberculin tests and positive histoplasmin skin tests.

Natural history is known.

CANDIDIASIS (MONILIASIS)

(Pulmonary code No 360-208)

(Thrush of Mouth code No 610 209)

Candidiasis is an infection of the mouth vagina skin and nails. It may rarely involve the lungs and meninges. The diagnosis of the pulmonary form may be difficult because of scanty purulent exudate common finding of hyphae appearance is similar to that of tuberculosis but tubercle bacilli cannot be demonstrated in the sputum. One must demonstrate the constant presence of *Candida albicans* before the diagnosis can be maintained. However the organism may occur as a normal inhabitant of the mouth so greater care must be taken in making the diagnosis of pulmonary moniliasis.

Treat the oral infection with alkaline mouth washes and topical application of gentian violet diluted 1:10,000 in 10 percent alcohol for 4-5 days. Treat vaginal infection with alkaline douches douches of potassium permanganate 1:1500 or gentian violet 1:10,000. Protonate vaginal jelly in yeast. For fungous infections soak in olive oil parts twice daily in 1:2000 potassium permanganate for 30 minutes. Follow with 1 percent gentian violet topically in 15 percent alcohol. For genital and anal infection nystatin (Mycostatin®) is an antifungal substance of value in eliminating or reducing the number of *Candida* in the stool. It is recommended as a prophylactic therapy in cases of patient on intensive antibiotic therapy with all antibiotics who may develop vaginal *Candida* infection. Give 500,000 unit tablet three times daily until it is all fungi suppressed and determined by culture of stools. Dose may be doubled if necessary and the drug may be given concomitantly with any of the common oral antibiotics. Continue nystatin throughout the course of therapy with the antibiotics oral and for short period after the administration of the antibiotics has been discontinued.

CRYPTOCOCCOSIS OR TORULOSIS

(Of Skin code No 110 21x) (Meningitis code No 910 21x)

Cryptococcus involves chiefly the central nervous system but may invade the structures. It is world wide in distribution and caused by *Cryptococcus neoformans* (*Torula histolytica*). The characteristic lesions are pustules granulomatous ulcers and nodules. Meningeal involvement is the usual central nervous

system lesion. The disease is usually mistaken for tuberculosis meningitis if the organisms are not found.

No specific treatment is available. Treat symptomatically.

IMMUNIZATION SCHEDULES

Biologicals for immunization purposes are gradually being modified and concentrated. The schedules below do not apply to all preparations; follow the manufacturer's instructions which accompany the preparation.

Children

- 1 During first year
 - a Uncombined method. Pertussis vaccine: three subcutaneous injections of 0.5 cc each at one month intervals beginning at 2 to 3 months of age. Diphtheria-tetanus toxoid (alum or aluminum hydroxide): subcutaneous injections of 0.5 ml and then 1.0 cc at one month intervals beginning at 6 months of age and smallpox vaccination at 6 months to 1 year of age. Repeat if a take does not occur.
 - or b Combined method. Diphtheria-pertussis-tetanus (combined): three subcutaneous injections of 0.5 cc each at one month intervals starting at 2 to 6 months of age and smallpox vaccination as with uncombined method.
- 2 At two years. Schick test and booster dose of diphtheria-pertussis-tetanus mixture: 0.5 cc by subcutaneous injection.
- 3 At school age. Repeat procedure as for two years and do vaccination (repeat if a take does not occur).
- 4 Smallpox vaccination every 5 to 7 years or on exposure.

Adults

Adults traveling to foreign countries should obtain a list of required immunizations when applying for passports. Those living in endemic areas should maintain their immunization.

- 1 Smallpox vaccination. Repeat every 5 years or on exposure.
- 2 Typhoid (or typhoid-paratyphoid) vaccine (1 billion organisms per cc): 0.5 ml and then 1.0 cc by subcutaneous injection at weekly intervals. Repeat series every 3 years.
- 3 Yellow fever vaccine (Africa-South America): 0.5 cc subcutaneous.
- 4 Typhus vaccine (Cox type) (Europe-Asia-Africa-Australia-South America and Mexico): 1.0 cc subcutaneous repeated 7 to 10 days later for total of 2 doses. Repeat 1.0 cc every 4 to 6 months.
- 5 Cholera vaccine (Asia-Near East-East Indies): 0.5 and then 1.0 cc every 4 to 6 months.
- 6 Plague vaccine (2 billion organisms/cc) (Egypt-Asia-East Indies): 0.5 and 1.0 cc subcutaneous at interval of 7 to 10 days.
- 7 Tetanus toxoid: 1.0 cc subcutaneous repeated at 30 and 60 days for total of 3 doses. Booster injection of 1.0 cc 1 year later and on injury.
- 8 Diphtheria immunization in adults who are Schick positive is frequently followed by severe local and general reactions. A Koloney test (0.1 cc of 1:20 dilution of fluid to skin) should be applied intradermally. If negative: 0.5 ml and

then 10 cc should may be given at monthly intervals. If positive inject intradermally 0.1 cc of 1:10 dilution of toxin at 3 to 4 week intervals for 3 doses.

HYPERSENSITIVITY TESTS AND DESENSITIZATION

Before injecting antitoxin or similar material derived from animal source always perform the following test for hypersensitivity. If both tests are negative no desensitization is necessary and a full dose of the antitoxin may be given. If one or both of the tests are positive desensitization is necessary.

A Intradermal Test. Inject 0.1 cc of a 1:10 dilution of the antitoxin intradermally into the skin of the flexor surface of the forearm. A positive test is manifested within 30 minutes by the occurrence of a large wheal and surrounding areol.

B Conjunctival Test. Instill 1 drop of a 1:10 dilution of the antitoxin into the conjunctival sac of one eye as a test dose and 1 drop of physiological saline in the other eye as a control. A positive test is indicated by conjunctival injection, itching and edema occurring within 30 minutes.

Desensitization

A Preliminary Measure

- 1 Antihistaminic drug should be administered before beginning desensitization in order to lessen any reaction that might occur (see p. 45).
- 2 Epinephrin U.S.P. Adrenalin B.P. 0.5 to 1.0 cc (8-15m) of a 1:1000 solution must be ready in a syringe for immediate administration.

B Desensitization Method. The following plan may be used in desensitization. Give doses 1 M at 30 minute intervals and observe closely for reactions.

- | | |
|---------------------------------|------------------------------|
| 1st dose 0.1 cc (1:10 dilution) | 7th dose 1.0 cc (undiluted) |
| 2nd dose 0.2 cc (1:10 dilution) | 8th and subsequent doses |
| 3rd dose 0.5 (1:10 dilution) | 1.0 (undiluted) every |
| 4th dose 0.1 cc (undiluted) | 30 minutes until the total |
| 5th dose 0.2 cc (undiluted) | amount of antitoxin is given |
| 6th dose 0.5 (undiluted) | |

Treatment of Reaction

- If mild reaction occurs stop back to the next lower dose and continue with the dose indicated. If a severe reaction occurs administer pinphrine as mentioned below discontinue the antitoxin until the test is negative again. Should doses not be hypotensive or titrated slowly using gradual increase of the antitoxin.
- If manifestation of a reaction appears give 0.5 to 1.0 cc (8-15m) of epinephrine subcutaneous. The symptoms include such as generalized edema dyspnea coughing hives or shock. Maintain a close observation of the patient and repeat pinphrine as necessary.

Chapter 20

ANTI INFECTIVE CHEMOTHERAPEUTIC AGENTS

Sulfonamides antibiotics aminosalicylic acid and isoniazid are used for the treatment of bacterial and rickettsial infections for the treatment or prevention of secondary bacterial infections in virus diseases and for prophylaxis against streptococcal infections in patients with valvular heart disease (to prevent subacute bacterial endocarditis)

Precautions in the Use of Chemotherapeutic Agents

- 1 *Etiologic diagnosis is of paramount importance*
- 2 Indiscriminate use may lead to serious toxic reactions
- 3 Insufficient dosage or unnecessary administration for minor illnesses may permit the emergence of resistant strains
- 4 Certain combinations may antagonize each other. Combined administration is probably best avoided except in exceptional clinical circumstances where the indications are clear
- 5 Topical administration (especially penicillin and Monomid) may sensitize the patient so that a severe hypersensitivity reaction may occur upon later systemic use

CHOICE OF ANTIBIOTICS IN BACTERIAL INFECTIONS

The choice of antimicrobial agents in the treatment of bacterial infections may be made in one of three ways

(1) The clinical appearance may be so characteristic of a given etiologic agent that specific antimicrobial therapy can be chosen without bacteriological examinations (E.g. meningococcal meningitis acute gonorrhea pneumococcal lobar pneumonia)

(2) The clinical appearance may be compatible with a variety of etiologic organisms in which case it is necessary to identify the specific organism by smear culture etc. When the organism has been identified the antimicrobial drug of choice can usually be selected on the basis of clinical experience (E.g. penicillin for streptococcal infections chloramphenicol for salmonella enteritis sulfonamides for meningococcal meningitis)

(3) If the drug of choice for an identified organism is not known (due to the variability of response to antibiotics on the part of some organisms e.g. staphylococci coliform bacteria) or if the organism itself is not known but can be isolated from clinical specimen antibiotic sensitivity test (see below) are required to determine which of several available antimicrobial agents is likely to have a bacteriostatic or bactericidal effect

Antibiotic Sensitivity Testing

The principles of antibiotic sensitivity testing are outlined below. However the immediate clinical situation must be borne

in mind and decide whether to wait for the results before proceeding with antimicrobial therapy. In most instances empirical therapy based on the assumption of the illogical assumption may be begun without necessity of test. In a very infectious disease it should be begun and later modified by sensitivity tests. It should be performed in bacterial diseases such as recurrent infection (especially of the urinary tract) and infections due to organisms likely to exhibit considerable strain variation in sensitivity.

A Plate Test is a late and hurried plate heavily with the clinical specimen (e.g., urine) through a baffle with a pour culture and wait for 48 hours until the plate is dry. Place small filter paper discs at intervals with various antibiotics on the plate 2-3 cm apart. Inubate overnight. Drugs which fail to give zones of inhibition are not likely to be clinically significant against the organism. NOTE: This is a crude rapid test which does not always correlate well with the results of the sensitivity test or with clinical response.

B Tube Test. The tube test measures more exactly the concentration for a antibiotic necessary to inhibit growth of a standard dose of organism under standard conditions. A series of broth tubes containing graduated amounts of an antibiotic is inoculated with a dilution of the broth culture of the test organism. After incubation the tubes are examined for turbidity. The end point is the last tube that contains no turbidity. The antibiotic used in the last tube remains clear. Upon the basis of a rough estimate of the volume of each tube to inhibit growth of the organism can be determined. In addition, bacterial identification may be determined by the tube dilution method.

SULFONAMIDE DRUGS

The sulfonamides are chemically related to sulfanilamide. New derivatives have wide antibacterial spectra and more effective pharmacological properties than the old sulfonamides. Since the activity of any sulfonamide compound may be predicted on the basis of certain physicochemical properties it is evident that maximal antibacterial effectiveness has been approximated by sulfadiazine, sulfamerazine, sulfamethazine and sulfisoxazole (Gastrin[®]). And the use of older sulfonamides is rarely if ever warranted. Sulfamethoxypyridazine (Kynex[®]) is slowly excreted and gives poor results as an effective agent.

Indications Antimicrobial Spectrum (Table 514)

The sulfonamides give a wide but still limited range of activity against pathogenic organisms. At the present time the sulfonamides are the only agents of choice only in meningococcal infection (Nissle and Ungitid[®]).

A Except for the infection mentioned below the sulfonamides should be used as alternate or additional agents to one of the antibiotics against infections of known susceptibility.

B Glucosulfone sodium (Promin[®]) this as well (Prominole[®]) and if necessary sodium (Diaso[®]) are more effective than the sulfonamides in how permeable within a limited area. Usually the addition of the drug against the following pathogens:

1. Mycobacterium tuberculosis
2. Mycobacterium leprae

ANTI INFECTIVE CHEMOTHERAPEUTIC AGENTS

Sulfonamides, tetracyclines, ampicillin, and isoniazid are used for the treatment of bacterial and rickettsial infections for the treatment or prevention of secondary bacterial infections in viral diseases and for prophylaxis against streptococcal infections in patients with valvular heart disease (to prevent subacute bacterial endocarditis).

Precautions in the Use of Chemotherapeutic Agents

1. Diagnosis of the infectious disease
2. Indications for use may lead to serious toxic reactions
3. Dosage or unnecessary administration for minor diseases may permit the emergence of resistant strains
4. Combination of drugs may antagonize each other. Co-based administration is probably best avoided except in exceptional cases where the indications are clear
5. Adverse reactions (especially penicillin and a Sulfonamide) may occur upon later systemic use

Choice of Antimicrobials in Bacterial Infections

The choice of antimicrobial agents in the treatment of bacterial infections may be made in one of three ways

- (1) The clinical appearance may be so characteristic of a given infection that a specific antimicrobial therapy can be chosen without laboratory examinations (E.g. meningococcal meningitis, streptococcal pneumonia)
- (2) The clinical appearance may be compatible with a variety of etiologic organisms in which case it is necessary to identify the organism by culture etc. When the organism has been identified the antimicrobial drug of choice can usually be selected on the basis of clinical experience (E.g. penicillin for streptococcal infections, chloramphenicol for salmonella enteritis, sulfonamides for meningococemia.)

(3) If the drug of choice for an identified organism is not known (due to the variability of response to antibiotics on the part of some organisms e.g. staphylococci, coliform bacilli) or if the organism itself is not known but can be isolated from clinical specimens, antibiotic sensitivity tests (see below) are required to determine which of several available antimicrobial agents is likely to have a bacteriostatic or bactericidal effect.

Antibiotic Sensitivity Tests

The principle of antibiotic sensitivity testing are outlined below. However, the immediate laboratory situation must be borne in mind.

C P t i s

- 1 Hemoglobin determination and white blood cell count every other day. Differential if WBC is less than 6000. Discontinue sulfonamid if granulocyte count is less than 50%.
- 2 Daily fresh urine for pH (use nitroblue paper) and estimate. Increase alkali (sodium bicarbonate) if pH is less than 7.0. Discontinue drug if red blood cells are found in urine (see above). Increase urine output if less than 1500 cc per day or crytilluria occurs (must be examined for in a fresh specimen).
- 3 Daily observation of patient for drug fever, rash, jaundice, nausea, vomiting, etc.

Contraindication to Sulfonamid

- 1 History of previous severe reaction
- 2 Renal insufficiency (Very small doses may be used with caution)
- 3 Liver damage (Proceed with caution if essential)
- 4 Heart failure (If sulfonamides are absolutely necessary, substitute potassium bicarbonate for sodium bicarbonate as alkalinizing agent)

AMINOSALICYLIC ACID (PAS)

Aminosalicylic acid (PAS) and its dimethyl salt have been found to exert considerable tuberculostatic activity. They are bacteriostatic to streptomycin in many susceptible strains of *M. tuberculosis*. The simultaneous administration of PAS and streptomycin in the treatment of strains of *M. tuberculosis* resistant to the latter. In addition to the bacteriostatic effect, antipyretic activity is present.

PAS is absorbed daily from the gastrointestinal tract. Peak serum concentration is reached in 30 to 60 minutes and minimum level is reached again in 4 hours. PAS may be administered in intravenous and subcutaneous solutions.

Dosage and Route of Administration

- A. Oral 3 to 4 Gm (45 to 60 g) as the acid every 6 hours.
 B. Intravenous 15 Gm in 3% solution given in 2 doses 4 hours apart. 5 mg of heparin should be added to each liter.

Toxicity

Nausea, vomiting, diarrhea, drug fever, dermatitis, erythema, hematuria, and hypoproteinememia may be observed. Gastrointestinal symptoms may possibly be avoided by parenteral administration of sodium PAS.

ISONIAZID (INH)

Isoniazid (INH) and related compounds possess considerable bacteriostatic activity. Cross resistance to streptomycin and PAS does not exist. Bacterial resistance to INH develops rapidly. INH is readily absorbed from the gastrointestinal tract and distributed throughout the body fluid, including the cerebrospinal fluid.

Oral Sulfonamides Adult and Pediatric Dosage Schedules

Indications & Preparations	Adult Dosage	Pediatric Dosage
Meningitis		
Initial dose: One of sulfonamide or sulfonamide mixt	2-4 Gm (50-60)	20 mg (4/3 gr) /lb body wt
Sustaining dose: 1/2 meq asin d m thod)	0.5 Gm (7 1/2 gr) 4 q 6 H	5 mg (4/12 gr) /lb 4 q 6 H
or 1 (Gantogen)	1 Gm (15 gr) 4 q 6 H	mg 4 q 6 H
or ampicillin 1/2 d in d 1/2 d in	1 Gm (15 gr) 4 q 6 H	mg 4 q 6 H
Sustaining dose	1 Gm (15 gr) 4 q 6 H	10 mg (1/6 gr) /lb 4 q 6 H
or Sustaining dose	1 Gm (15 gr) 4 q 6 H	10 mg (1/6 gr) /lb 4 q 6 H
or Sustaining dose	1 Gm (15 gr) 4 q 6 H	10 mg (1/6 gr) /lb 4 q 6 H
(Myxer)	1 Gm (15 gr) 4 q 6 H	10 mg (1/6 gr) /lb 4 q 6 H
Urinary tract infection: On sulfonamide mixt	0.5-1 Gm (7 1/2-15 gr) 4 q 6 H	5-10 mg (1/12-1/6 gr) /lb 4 q 6 H
Prophylaxis: 1/2 d in 1/2 d in	0.5 Gm (7 1/2 gr) b i d	5 mg (1/12 gr) /lb b i d
It is important to Sulfaguanidine	50 mg (3/4 gr) /lb S t t h	25 mg (3/8 gr) /lb 4 q 6 H
Leprosy: 1/2 d in 1/2 d in	4 (1 d) 4 q 6 H	0.5-1 (1/2-3/4 d) 4 q 6 H
Thi... (P...)	1-2 Gm (15-30 gr) 4 q 6 H	0.25-1 Gm (1/4-1 gr) 4 q 6 H
Orally... (gl...)		

Toxicity and Management

A. Toxic Reactions

1. Mild: Continue therapy if necessary. Symptoms include nausea, vomiting, headache, dizziness, crystalluria. Moderate: Stop therapy until continuation is essential to life. Symptoms include fever, stomatitis, conjunctivitis, rhinitis, diarrhea, microhematuria, psychosis.
3. Severe: Stop therapy and give fluids. Symptoms include granulocytopenia, hemolytic anemia, aplastic anemia, thrombocytopenia, hypotension, exfoliative dermatitis, severe hepatomegaly, oliguria, leukopenia, etc.

B. Allergic Reaction A considerable percentage of individuals who have previously received sulfonamides develop allergic reactions more than 7 days become sensitized and may develop immediate and severe reaction on administration. Fever, angioedema, urticaria, and other rash and pruritus may occur.

History of previous administration should be obtained. Cross sensitivity to various sulfonamides may exist. Severe symptoms may be avoided by giving a test dose of 0.5 Gm (7 1/2 gr) and observing for 6 hours.

toyed Acquired penicillin resistance is not commonly countered clinically

Absorption Distribution, Elimination

- A Absorption** Penicillin is well absorbed rapidly absorbed when administered intravenously or intramuscularly and somewhat more slowly absorbed after subcutaneous injection. The peak concentration in the blood is reached immediately after intravenous injection and within one hour after intramuscular injection. Blood levels peak at 2 to 3 hours after doses of 1 as the 50 000 units intramuscularly and somewhat longer with larger doses. Penicillin procaine suspension is a poor mess absorbable concentration is for 12 to 48 hours and for 96 hours when mixed with 2% aluminum monostearate. Benzathine penicillin may produce measurable concentration for months after injection of 600 000 to 1 200 000 units. With liposoluble form maximum serum concentration tends to be lower than with aqueous solution. Dose does not appear to be related to high serum concentration. Penicillin is not absorbed from the stomach. It is absorbed readily from the small intestine. Approximately 5 times the intramuscular dose must be given to produce comparable blood levels. Antacids and buffer should be avoided because they decrease gastric juice absorption. It is best taken with an empty stomach. Penicillin is not destroyed by gastric acid. Penicillin is poorly absorbed from the stomach and instantly absorbed from the small intestine. The concentration of penicillin in serum and other body fluids may be measured by various biochemical methods.
- B Distribution** Penicillin is distributed throughout the body fluids but penetrates the joints pleural peritoneum and subarachnoid spaces irregularly. Penetration is more likely to occur if inflammation exists. Penicillin penetrates into the cerebrospinal fluid slowly after it has disappeared from the blood hence not sublethal levels are necessary most infection. Organism exposed to penicillin does not multiply for several times after exposure.
- C Elimination** Penicillin is excreted principally through 80% of the urinary tract in tubules and may be partially blocked by cholestyramine and probenecid. Penicillin is also excreted in the bile (Dodd¹⁹) and in the sweat (Beck²⁰).

Preparations

- A Commercially Available Preparations**
1. Crystalline penicillin (sodium potassium salts)
 2. Penicillin procaine oil
 3. Penicillin procaine oil with 2% methylmagnesium stearate
 4. Penicillin procaine aqueous suspension (may be combined with crystalline penicillin sodium)
 5. Penicillin tablets with or without buffer and benzathine penicillin (50 000 to 200 000 unit penicillin) and benzathine penicillin (60 000 units penicillin) of 100 000 units penicillin
 6. Penicillin V tablets 125 mg of 250 mg
 7. Penicillin powder for insufflation (50 000 unit penicillin a tridg)
 8. Penicillin oil emulsion in vegetable oil (general diphthylmethyl) (500 2000 nit penicillin gm)

Dosage and Routes of Administration

Penicillin G 5 to 10 mg (1/12 to 1/6 gr)/Kg body weight per day in 2 or 3 doses orally. Ten mg (1/6 gr)/Kg should be given daily in tuberculous meningitis. Sterile solutions may be given I.M.

Toxicity

Constipation, dysuria, hyperreflexia, postural hypotension and dizziness, eosinophilia, slight anemia, occasional casts and traces of albumin in the urine, reducing substances in the urine.

PENICILLIN

Penicillin is prepared from the cultural products of the molds *Penicillium notatum* and *Penicillium chrysogenum*. The commercially available preparations are crystalline sodium, calcium, potassium, and procaine salts of penicilloic acid.

Many types of penicillin, F, G, O, V, X, and K, are produced by the mold. Commercial penicillin is principally penicillin G, Penicillin X, which occurs only in small amounts, exhibits a slightly different range of antibacterial activity. Penicillin K becomes bound to serum protein and is relatively inactive therapeutically.

The Oxford and International units of penicillin are measured in comparison to the bacterial inhibitory power of a standard penicillin. Crystalline sodium penicillin contains approximately 1500 units per milligram. Dried crystalline penicillin retains its potency indefinitely, but watery solutions may deteriorate especially when not refrigerated.

Indications and Antimicrobial Spectrum (See table, p. 514)

Penicillin exerts bacteriostatic and bactericidal activity against a wide variety of pathogenic agents, but the susceptibility of these agents to penicillin may vary considerably. Clinical response of infections may be predicted with fair accuracy by means of in vitro sensitivity tests of the infecting organism. The procedure should be performed when expected therapeutic response does not occur or in the case of infections due to organisms such as staphylococci or *Streptococcus fecalis*, many strains of which are naturally resistant to penicillin.

Penicillin is indicated when infection with an organism known to be generally susceptible is diagnosed. Pneumonia, Rheumatic fever, specific infection, not a disease, e.g., pneumococcal pneumonia, not pneumonia, streptococcal pharyngitis, not acute pharyngitis. For specific indications, see under diseases in question.

Mode of Action, Resistance

Penicillin is both bacteriostatic and bactericidal. Its exact mode of action is not known, but in some way it probably interferes with the reproductive process of the organism.

Certain organisms produce penicillinase, which inhibits penicillin's activity. This may occur intracellularly, as in the case of *E. coli*, and some strains of staphylococci. Susceptible organisms exist in a lethal concentration of penicillin may acquire resistance. The mutants of the original organism which we encounter rarely resistant survive and multiply while the susceptible organisms are being

penicillin dissolved in 10 cc of physiologic saline should be administered 6 cc a day until the cerebrospinal fluid glucose content becomes normal. Penicillin should also be given intramuscularly.

- 3 Int pral intra artula 10 000 to 200 000 unit of p m illn may be introd ed into joint o pl ral paces in fact d by usc ptibl organism daily o every other d y f llowing aspiration
- 4 Oral Troch of p nicillin may be dissolv d slowly in the mo th in the t stime t of Vinc nt s stomatitis and pharyn gitis This form of therapy is vslu l s in oth forms f pharyngitis and may p od c stomatitis
- 5 Wounds and skin Solut ons of p nicillin c ntaining 200 1000 units per cc m y be ed s a w t dre ing in infect d wounds Peni illin is of no val e as n i gating solution because of the neces sity of prol nged conta t to prod ce a ttra terial off t

Contaminants from the air incorporated during use may be a danger in infections of the skin due to a septible organ form.

Twelve

Some of the principal features of penicillin therapy are almost unknown. Selection may be pre-extinguished. Fever and shock especially uterine may appear during the course of penicillin administration as long as a very low dose is used. The extremely minute cumulative dose. True idiosyncrasy to penicillin is rare. Immunity is largely limited to individual sensitivity. Inadequate phytochemicals. Potentially intradermal test to weak penicillin reactions may be based. Decontamination by attempt. Potentially to be sensitive to penicillin may be treated with penicillin. Oral penicillin G which may be substituted for aqueous penicillin. Expectation in the same dose. Cases of sensitivity occur locally and should be guarded against.

STREPTOMYCIN

St ept my is p pa d from the ult i p od cts of St epto
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1 W k m n unit d l Gm equals 1 000 000 Waksman unit

1 dicatlo a d A umle ob(a) Sp tr m (S table p 514)

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The indication for stentomy is limited to the following conditions:

B Strength of Solutions and Suspensions

- 1 Crystalline penicillin is generally given in a concentration of 10 000 units per cc but may be much more concentrated
- 2 Penicillin for subarachnoid injection should not be more concentrated than 1000 units per cc
- 3 Penicillin procaine complex in aqueous suspension may be prepared in concentration of 300 000 to 1 200 000 units per cc

Dosage and Methods of Administration

- A Intermittent Intramuscular. Penicillin in aqueous solution may be given in doses of 5000 to several million units every 3 hours intramuscularly. This remains the method of choice in most severe acute infections. In many infections equally good results may be obtained by administration of 100 000 to 300 000 units every 12 hours intramuscularly. Intramuscular injection of 300 000 to 600 000 units of penicillin procaine may be given every 24 hours. Penicillin procaine in oil with 2% aluminum monostearate produces measurable blood levels which may persist as long as 96 hours. Benzathine penicillin 600 000 to 1 200 000 units produces measurable serum concentrations for one month and is ideally suited to prophylactic use. These preparations are highly satisfactory except in the most severe acute infections.
- B Continuous Intramuscular and Continuous Intravenous. Where very high doses of penicillin are necessarily in the treatment of infections due to resistant organisms administration by continuous drip is often advantageous. Many million units dissolved in 1000 to 2000 cc of physiological saline or 5% glucose solutions may be given by indwelling needle or catheter in 24 hours. The intramuscular route should be changed as frequently as irritation occurs. Thrombophlebitis as a complication of intravenous administration may be avoided by hanging the vein used by addition of 10 mg heparin sodium to the solution.
- C Oral. Penicillin may be given orally in all but the severest of infections or oral medication may be substituted for parenteral after initial response to treatment. Doses of 100 000 units every 3 hours to 300 000 units every 6 to 8 hours may be given. Penicillin V may be given in a dose of 125 to 250 mg every 8 hours.
- D Topical
- 1 A solution 50 000 to 100 000 units may be aerosolized from 3 to 8 times a day. A solution containing 50 000 units per 0.5 cc may be nebulized by means of Vapo-phrin® or DeVilbiss No 40® nebulizers. Forced deep inhalation followed by retention of the inspired penicillin in gas possible should be insured. Hand pumping or compressed gas fed through a Y tube may be used to nebulize the solution. While local effect in the respiratory passages for the treatment of bronchitis and chronic bronchitis is usually the objective appropriate blood concentrations of penicillin frequently result. Secondary infection occurs commonly.
 - 2 Intrathecal. Although penicillin may penetrate the subarachnoid space after intramuscular injection this phenomenon is inconstant and may be delayed. Therefore in meningitis due to susceptible organisms 10 000 units of

Toxicity

Painful local reaction is uncommon at places. Drug rash is of many sort occur drug fever may be observed and slight nausea and dizziness are frequent. Eosinophilia may be noted but appears to have no significance. Cylindrocapsa and nitroglycerin retention not associated with permanent renal damage have been reported. Vestibular damage often manifested first by tinnitus and characterized by severe vertigo and ataxia follows high or prolonged dosage. If streptomycin is discontinued immediately recovery usually follows. Nystagmus and damage to bones permanent at all stages compensation is usually made by the patient. Deafness also may occur but it is rare. Vestibular apparatus disorder is less common with dihydrotetracycline but deafness may develop after treatment has been stopped. The same combined streptomycin and dihydrotetracycline in equal parts reduces the incidence of deafness and vestibular damage. Pleoptyosis in the serous protein content of the spinal fluid a bacterial block or myelitis may follow prolonged intrathecal administration of streptomycin.

TETRACYCLINE GROUP (CHLORTETRACYCLINE OXYTETRACYCLINE TETRACYCLINE)

The chemically related groups are similar antimicrobial spectra and pharmacological properties. Organisms resistant to one drug are usually resistant to the others although significant variation occasionally occurs. Generally they are highly inter interchangeable.

CHLORTETRACYCLINE (AUREOMYCIN[®])

Chlortetracycline (Aureomycin[®]) is prepared from Streptomyces ofaci. It is available as the hydrochloride.

Indication and Antimicrobial Spectrum (See table p. 514.)

Chlortetracycline (Aureomycin[®]) is an alternative to penicillin in the treatment of most bacterial infections. It is a broad spectrum antibiotic with a wide therapeutic range. It is active against most gram-negative and gram-positive cocci, the spirochetes of leptospirosis, rickettsiae of typhus, Rocky Mountain spotted fever, scrub typhus, Q fever and rickettsialpox. It is highly effective against the virus of lymphoplasma pneumoniae and the primary type of pneumonia.

Absorption and Elimination

Chlortetracycline is absorbed slowly from the gastrointestinal tract and peak blood concentrations are reached in from 2 to 4 hours depending on the dose and the rate of administration. Following administration of a single dose, high blood concentrations are maintained for a period of 6 to 24 hours, varying with the dose. Chlortetracycline is poorly absorbed after mucous membrane and subcutaneous administration. Accumulation occurs in the

stituting treatment. Most tubercle bacilli become streptomycin resistant within 3 months of the beginning of treatment although the simultaneous use of PAS or isoniazid may delay this event. A dose of one or both should always be used with streptomycin in tuberculosis.

Mode of Action Resistance

Streptomycin is both bacteriostatic and bactericidal. Its mode of action is unknown. Resistant variants of organisms may multiply quickly in infections treated with streptomycin so that further therapy with the antibiotic is useless. Streptomycin should be used only when necessary and adequate initial dosage should be used to prevent development of drug resistance.

Absorption Distribution Excretion

- A Absorption Streptomycin is readily absorbed from the site of intramuscular injection. The peak serum concentration is reached within one hour and detectable amounts are present up to 6 hours later. It is likely that streptomycin persists longer than this in the tissues. If streptomycin is administered every 3 to 4 hours gradually increasing serum levels will be noted due to slow accumulation. Administration every 6 hours is sufficient in all but the most severe infections in which cases the drug should be given initially every 3 or 4 hours. Streptomycin is not absorbed from the gastrointestinal tract but exerts bacteriostatic activity in the lumen of the bowel.
- B Distribution Streptomycin is distributed throughout the body similarly to penicillin. Penetration of the cerebrospinal fluid is inconstant and unreliable.
- C Excretion Streptomycin is excreted principally in the urine where the concentration exceeds that of the serum.

Dosage and Routes of Administration

- A Non-tuberculous Infections 1 to 5 Gm. daily may be given intramuscularly in divided doses every 3 to 6 hours. Most acute generalized infections require approximately 2 to 4 Gm. per day. Urinary tract infections due to highly susceptible organisms may be treated with 500 mg. intramuscularly every 6 hours for 5 days. Streptomycin should not be used in the presence of obstruction of the urinary tract because of the necroticity of the development of resistant organisms.
- B Meningitis Intrathecal intramuscular administration 25 to 50 mg. dissolved in 10 cc. of physiological saline solution may be given intrathecally once daily until the cerebrospinal fluid sugar content becomes normal.
- C Bacillary Dysentery Streptomycin may be given orally 0.5 Gm. (7 1/2 gr.) every 6 hours for 5 to 7 days.
- D Tuberculosis 0.5 Gm. of streptomycin and 0.5 Gm. of dihydrotetracycline intramuscularly twice weekly is indicated in non-disseminated forms of tuberculosis. In acute cases of tuberculous pneumonia and milary tuberculosis 40 mg./Kg. of body wt. (20 mg./lb.) daily should be given. In tuberculous meningitis 60 mg./Kg. of body wt. (30 mg./lb.) daily should be administered intramuscularly in addition to 2 mg./Kg. of body wt. (1 mg./lb.) per day intrathecally until isoniazid is used simultaneously. (See Tuberculous Meningitis p. 468.)

Contraindications The preparation for I.M. use may be given in a dose of 0.5 mg. every 24 hours or 0.1 Gm. every 6 hours

Toxicity

Nausea vomiting diarrhoea stomatitis and dermatitis occur occasionally. Hepatitis may result from prolonged intravenous treatment at high dosage. Thrombophlebitis may result from intravenous administration. Superinfection with resistant staphylococci may occur usually as severe enterocolitis. This also occurs with other broad spectrum antibiotics.

TETRACYCLINE (ACHROMYCIN® TETRACYN®) POLYCYCLINE® STECLIN® PANMYCIN®)

Tetracycline is produced by removing the chlorine from chlorotetracycline. It is similar to chlorotetracycline and oxytetracycline but is more soluble in solution than either derivative.

Indications and Antimicrobial Spectrum (See table on p. 514)

Tetracycline is a broad spectrum antibiotic whose field of activity is similar to those of chlorotetracycline and oxytetracycline. Susceptibility of strains of bacteria may differ among the three drugs however.

Absorption and Excretion

Tetracycline is absorbed and excreted similarly to chlorotetracycline. It may diffuse more readily into the cerebrospinal fluid. Phosphate buffer gastric secretion may alter the rate of absorption.

Dosage and Route of Administration

- A Oral 0.25 to 1.0 Gm. every 6 hours
- B Intravenous 0.5 to 1.0 Gm. every 12 hours
- C Intramuscular 0.1 Gm. every 8-12 hours

Toxicity

Similar to that of chlorotetracycline and oxytetracycline but significantly less frequent.

CHLORAMPHENICOL (CHLOROMYCETIN®)

Chloramphenicol (Chloromycin®) originally prepared from Streptomyces aureofaciens is produced synthetically.

Indications and Antimicrobial Spectrum (See table on p. 514)

Chloramphenicol is a broad spectrum antibiotic with a wide range of activity. It is indicated for the treatment of lymphogranuloma venereum, psittacosis, diphtheria, typhoid pneumonia. Generally speaking it is more effective than chlorotetracycline and oxytetracycline in typhoid fever. Particularly useful in effect against other gram-negative organisms especially the enteric bacilli. Many staphylococci are susceptible to chloramphenicol.

Absorption and Excretion

Chloramphenicol is rapidly absorbed from the gastrointestinal tract.

body at high dosage so that blood levels become increasingly elevated during prolonged administration at high dosage. Chlorotetracycline is excreted slowly by the kidney. It does not appear daily in the cerebrospinal fluid or pleural fluid, but it is present in high concentration in the urine and stools.

Dosage

- A Oral 0.25 to 1.0 Gm. may be given orally every 6 hours.
 B Intravenous. Similar results may be obtained by the intravenous administration of 100 mg. every 6 to 8 hours or 500 mg. every 12 hours. In resistant infections combined oral and intravenous therapy may be used.
 C Intramuscular. 250 mg. in 1% procaine solution with 50 units of hyaluronidase added every 6 hours may be substituted for intravenous therapy when required.

Method of Administration

250 mg. orally every 6 hours appears adequate in most acute infections. Gastrointestinal symptoms may be minimized by administering the drug only when food is in the stomach or by simultaneously administering carboxymethylcellulose. Superinfection with yeast in the oropharynx and perineal area may occur but are probably secondary infections of local sensitivity reactions.

Toxicity

Nausea and vomiting are common following oral administration but this may be avoided by reducing the dose to 250 mg. every 6 hours or administering the drug intravenously. Rashes and stomatitis may occur. Loose bowel movements may be observed.

OXYTETRACYCLINE (TERRAMYCIN®)

Oxytetracycline (Terramycin®) is derived from Streptomyces rimosus. The commercial preparations are the hydrochloride and the base.

Indications and Antimicrobial Spectrum (See table on p. 514)

Oxytetracycline is a broad spectrum antibiotic whose range of activity is similar to that of chlorotetracycline. It may be used in infections due to gram positive and gram negative cocci, gram positive and gram negative rods, pleuropneumoniae, tetrads and the viruses of primary atypical pneumonia, lymphoplasma venereum and pinta.

Absorption and Excretion

Oxytetracycline is completely absorbed from the gastrointestinal tract. Satisfactory serum levels may be maintained by administration every 6 hours. Excretion is principally by the kidneys. Significant amounts appear in the bile. Appearance in the cerebrospinal fluid is delayed and irregular.

Dosage and Routes of Administration

- A Oral 0.25 to 1.0 Gm. may be given orally every 6 hours.
 B Intravenous 0.5 to 1.0 Gm. may be administered every 12 hours.
 C Intramuscular Oral therapy should be used whenever possible.

T xic tv

Mild toxic effects occur at dosage levels over 2.5 mg/Kg of body wt./day. Adverse reactions are usually reversible. Widespread drowsiness, ataxia, numbness of the fingers and feet, impairment of vision, blurring of the vision, diplopia and vertigines may occur. Allergic reactions such as tingling, erythema, wheezing, dyspnea have been observed. Irritation at the site of intramuscular injection is common.

BACITRACIN

Bacteria is derived from the growth products of *Bacillus subtilis*.

Ind. tit. and A. tit. of al. Sp. trum. (See table on p. 514.)

Ba it can is tiv ga nst gram po tiv occi and p och tes
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Abstract and Introduction

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510 Polymyxin

tract reaching a peak serum concentration within 2 hours. Absorption following rectal administration is slightly less efficient. 0.5 Gm. may be administered intramuscularly or intravenously every 6 hours. Excretion is principally by the kidneys and high concentrations are reached in the urine.

Dosage and Routes of Administration

- A Oral Adult 0.5 Gm. every 6 h. s. children 40 mg./Kg./day
- B Rectal 125 to 150 mg./kg. (55 to 70 mg./lb.) per day in children every 6 hours. Capsule should be punctured before insertion
- C Intramuscular and Intravenous 500 mg. every 6 hours

Toxicity

Nausea and vomiting, diarrhea, nervous depression, dermatitis, granulocytopenia and aplastic anemia occur occasionally. Therefore chloramphenicol should be used only on definite indication.

TYROTHRICIN

Tyrothricin is prepared from *Bacillus brevis*. It is used topically as an ointment or watery suspension. It is active only against gram positive organisms. Because of toxic effects on parotid administration, its use is limited entirely to the topical treatment of infected wounds and pyoderma.

POLYMYXIN (AEROSPORIN®)

The polymyxins of which B, D and E have been given clinical trial are derived from *Bacillus polymyxa* and related organisms.

Indication and Antimicrobial Spectrum (See table on p. 514)

With the exception of most strains of *Proteus vulgaris*, polymyxin is bactericidal against gram negative rods and most strains of *Pseudomonas aeruginosa* (pyocyanins). Polymyxin is indicated in severe systemic infections due to gram negative rods particularly infections due to *Pseudomonas aeruginosa* which do not respond to other forms of chemotherapy. It may be used as a local application in wounds infected with susceptible organisms. It may be given orally in the treatment of the sigmoiditis after attack.

Absorption and Excretion

Absorption is rapid following intramuscular injection. Excretion is largely by the kidney and high concentrations are achieved in the urine. Polymyxin is not absorbed from the gastrointestinal tract and when it is given by mouth it has its principal activity in the lumen of the bowel.

Dosage

- A Intramuscular 1.5 to 2.5 mg./Kg. of body wt. divided 4 to 3 or 4 doses
- B Oral 20 mg./Kg. of body wt./day given in 3 or 4 doses

ties in as n and b tereml d not exist Toxic re ctions in clud gastrointestinal irritat n and occas al skin rashes

Dose Adults 100 mg or lly 4 tim s daily Child en 5 to 8 mg /Kg /day Anti tra o s preparation is now available but its at lin ind tio a ot y t know

VIOMYCIN (VINACTANE® VIOCIN®)

V my i is de iv d fr m St ptomy puni eus It is active only gainst My ob cte lum tuberc l is in luding train esis tant t st ept mycin aminosal ylc a d and i oniazid Because it is highly n phr toxic and neurot xic t use is very limited Tx r action in lud eighth n ve dam ge and enal in uff cncy with di tu bed lect olyte balan e

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NYSTATIN (MYCOSTATIN®)

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Dose O ally 500 000 units 3 times daily Locally as vagl al ppo it l s (100 000 nt) on o twied ly or s tm t (100 000 unit /Gm)

NOVOBIOCIN (ALBAMYCIN® CATHOMYCIN®)

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OLEANDOMYCIN (MATROMYCIN®)

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Dose 0.25 Gm v ry 6 ho

512 Nitrofurantoin

mycin is principally excreted by the kidney and appears in the urine in high concentration

Dosage

- A Topical Ointments containing 1000 units per gram or solutions contain g 200 units per cc may be used locally
- B Oral 0.1 Gm /Kg daily divided into 4-6 doses
- C Intramuscular 15-20 mg /Kg daily divided into 4 doses

Toxicity

Renal damage manifested by albuminuria. Nephrogenic reaction may occur. Deafness may follow parenteral administration

ERYTHROMYCIN (ERYTHROCIN® ILOTYCIN®)

Erythromycin is a medium spectrum antibiotic derived from *Streptomyces erythreus*. Its action may be bactericidal or bacteriostatic depending on the susceptibility of the bacteria. Resistance to erythromycin may develop rapidly under certain circumstances most notably by staphylococci. For this reason erythromycin should not be used alone in serious staphylococcal infections.

Indications and Antimicrobial Spectrum (See table on p. 514)

Erythromycin is active against most strains of gram positive cocci, gram negative cocci, *C. diphtheriae*, *H. influenzae*, *H. pertussis* and *B. illae*. Activity has also been shown against the viruses of lymphopathia venereum and psittacosis and the rickettsias of typhus. Erythromycin may be used in infections due to these organisms as an alternative to penicillin and other antibiotics.

Route of Administration and Dosage

- A Oral 0.2 to 0.5 Gm every 6 hours
- B Intravenous 0.5 Gm every 12 hours

Toxicity

Nausea, vomiting and diarrhoea occur occasionally.

FUMAGILLIN (FUMIDIL®)

Fumagillin is derived from *Aspergillus fumigatus* H-3. It is directly amebicidal and apparently is effective also against other enteric protozoa. It has been valuable in the treatment of drug-resistant amebiasis. Dosage 30 to 60 mg orally daily for 10 days.

NITROFURANTOIN (FURADANTIN®)

Nitrofurantoin is active against a wide variety of bacteria, both gram positive and gram negative. It is readily absorbed from the gastrointestinal tract and excreted in high concentration in the urine. Serum and tissue concentrations are insignificant. It is useful in the treatment of infection of the urinary tract when significant

Chapter 21

DISEASES OF UNKNOWN ETIOLOGY

A variety of names (collagen diseases, diffuse vascular diseases, vasculitis, etc.) have been given to a group of diseases which appear to have in common a pathological involvement of mesenchymal tissues. Rheumatoid arthritis, disseminated lupus erythematosus, periarteritis nodosa, scleroderma, dermatomyositis, sarcoidosis (a sarcoidosis) and perhaps glomerulonephritis are the chief members of this group of rather ill-defined but probably related diseases of unknown etiology. The differentiation of these disorders is sometimes possible on clinical grounds, and in many instances the diagnosis can be established only after long continued and painstaking observation (see page 520). There is some evidence that hypersensitivity is common to many of these diseases although the pathological action in these cannot be traced probably caused by a wide variety of injurious agents.

Clinical Findings

Certain clinical features are common to many of the collagen diseases although they may be considered individually in their relative frequency and frequency of manifestation.

A Clinical Lesions But they are heterogeneous multiform and diverse.

B Rheumatoid Phenomena Purpuric hemorrhagic

C Arthritis Manifestation Synovitis, arthritis, arthralgia

D Cutaneous Manifestation Purpuric dermatitis, erythema, nodules, etc.

E Vascular Manifestation Arteritis, hypertension, etc. Renal disease, peripheral neuropathy.

F Lymphatic Involvement Lymphadenopathy and splenomegaly.

G Skeletal Manifestation Polyosteoarthralgia, periostitis, etc.

H Neuromuscular Manifestation

I Immune Manifestation Anemia, leukocytosis, leukopenia.

J Endocrine Manifestation

K Systemic Constitutional Manifestation Fever, weight loss, fatigue.

Laboratory Findings

The following list of special diagnostic significance.

A Skin Biopsy Characteristic histological changes may be observed in most of the conditions.

B Mucous Membrane Biopsy Characteristic histological changes occur in the oral cavity, etc.

C Lupus Erythematosus Clinical manifestations. The characteristic

ANTIMICROBIAL SPECTRA OF CHEMOTHERAPEUTIC AGENTS

Data representatively based on available literature and clinical experience. The degree of hold time may be prolonged further experience.

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$$\frac{1}{45} - \frac{1}{610} \ln 4 \approx C$$

a Sodium Salicylate U S P is most widely used

(1) Dose 12 Gm (15-30 g) very 2-4 hours orally

The drug should be given in sufficient doses to allay symptoms and fever and it may be necessary to give maximum dose to achieve this result. In an occasional patient maximum doses may not be completely effective. There is no evidence that intravenous administration has any advantage over the oral route.

(2) Toxic reactions. The early reactions include tinnitus, nausea and vomiting. Sodium salicylate may be given enterically at 0.5 Gm (7½ gr) pills or with equal doses of sodium bicarbonate to reduce gastric irritation. *Never use sodium salicylate or sodium bicarbonate in patients with acute rheumatic fever who have associated cardiac failure.*

b Acetylsalicylic Acid U S P may be substituted for sodium salicylate with the same dosages and precautions.

c Aminopyrine N F (Pyramido®) If the salicylate is not tolerated this drug may be used in doses of 0.2-0.4 Gm (3-6 gr) every 3-4 hours. Check the WBC very 2-4 days when giving this drug.

2 The sulfonamides and penicillin should never be used in the treatment of acute rheumatic fever; they are of no value and the sulfonamides may be harmful.

Prevention of relapse - There is some evidence that penicillin may prevent a relapse if used within 12-24 hours after the onset of a streptococcal infection.

b Coxsack infection. If an acute infection occurs during an attack of rheumatic fever, give antibiotic agents indicated (see page 514) but avoid giving sulfonamides.

B. General Measures

1 Bed rest should be enforced until all signs of active rheumatism have disappeared. The criteria for this are:

a Return of the temperature to normal with patient at bed rest and without medication.

b Normal sedimentation rate.

c Normal resting pulse rate (under 100 in adults).

d Return of ECG to normal (fixation of abnormality).

2 Gradual resumption of activities. Patient may be allowed up slowly but several months should elapse before return to full activity unless the infection was exceedingly mild.

3 Maintaining good nutrition.

C. Corticosteroids (ACTH) and the Cortisone. Although rather remarkable results have been observed in acute human infection, treatment with these drugs which improve manifestations only temporarily. They may be applied in the case of fever, malaise, toxicity and polyarthritides. Abnormal ECG changes (prolonged P-R interval) and blood sedimentation rate may return to normal limits within a week. Optimal dosage should be influenced by the drugs on the development of tuberculous and fungal infections have not been established (See page 423).

D. Treatment of Complications

1 Congestive failure. Treat as for congestive failure (see page 182) with the following cautions:

polymorphonuclear leukocytes (with round vacuole partially filled with nuclear material) in marrow and peripheral blood smears suggest disseminated lupus.

- D Antistreptolysin Titer Demonstration of changing A S titer may provide etiologic information regarding background of previous streptococcal infection which is of especial value in the diagnosis of rheumatic fever.

- E C reactive Protein (CRP) The test for this abnormal serum protein is said to provide a sensitive index of inflammatory activity.

Rheumatic fever and Sydenham's chorea will be discussed individually. The treatment of other diffuse collagen diseases will be discussed collectively. Acrocyanosis (acrosclerosis) has been included in this group for purposes of differential diagnosis although it is doubtful that it is a true member of the collagen disease group. Rheumatoid arthritis has been dealt with in Chapter 12 (see page 311) glomerulonephritis in Chapter 11 (see page 293).

RHEUMATIC FEVER (code No 010 932)

A generalized disease of unknown etiology usually coming on 1-3 weeks after an acute infection with the hemolytic streptococci and manifested usually by pathological changes involving the heart, blood vessels and serous surfaces primarily the joints. It has a marked tendency to recur.

Diagnosis

The diagnosis of active rheumatic fever can usually be made if the patient has 2 major manifestations or 1 major and 2 minor manifestations of the disease (Jones).

A Major Manifestations

- | | |
|----------------------------------------------------|--------------------------|
| 1 Definite pathologic of rheumatic fever | 3 Inflammation of joints |
| 2 Signs of active carditis (including ECG changes) | 4 Subcutaneous nodules |
| | 5 Chorea |

B Minor Manifestation

- | | |
|-----------------------|----------------------------|
| 1 Fever | 5 Non-traumatic nose bleed |
| 2 Erythema multiforme | 6 Purpura |
| 3 Abdominal pain | 7 Pneumonitis |
| 4 Pre-ordial pain | |

- C Laboratory Tests May show increased sedimentation rate, increased antistreptolysin titer, C reactive protein, leukocytosis and anemia and abnormal ECG.

Treatment (See page 187 for treatment of rheumatic heart disease)

A Specific Measures

- 1 Salicylate therapy The salicylate markedly reduce fever, alleviate joint pain and possibly reduce joint swelling. There is no evidence that they have any effect on the natural course of the disease. The salicylates should be continued as long as necessary to relieve pain, swelling or fever. If withdrawal of the drug results in a recurrence of symptoms they should immediately be reinstituted.

most attacks of acute rheumatism. If this proves to be prompt and efficient treatment of any preexisting infection in susceptible individual may be adequate prophylaxis.

CHOREA (Sydenham's) (code No 930 196)

Ales commonly manifest signs of rheumatic disease characteristically by generalized edema and congestion of the base and by involvement of the basal ganglia with vascular thrombosis. The hemorrhages perivascular infiltration and hemolysis of the cells. It occurs most frequently in females in the second decade and is characterized by jerky restlessness, effeminate mannerisms and tremors by dysarthria.

Treatment

- A Spinal Muscles Norkown Corticotropin (ACTH) and the response to be followed in many cases of the. It must be given relatively high initial dosage and marked sedation stimulation must be employed.
- B Febrile may be employed if all else fails. This may be given by either of 2 methods. Hypothalamic apparatus with temperature 39.5-40.5 C (103-105 F) for 3-5 hours twice weekly for 6-10 treatment. Typhoid fever is eliminated by 5-7 days.
- C General Measures
 1. General symptomatic measures and good nursing. If most important.
 2. Satisfactory with phenobarbital U.S.P. 15-30 mg (1/4-1/2 g) tid qid may be helpful.
 3. If valuable in the appropriate. Magnesium Sulfate U.S.P. 4-10 g (1 1/2 dr) of 10% solution I.M. or I.V. may be used. When administering magnesium sulfate I.V. slowly because a very fatal dose with 10% of 10% but a dose of 6 g of glomerulonephritis. Calcium is easy to administer I.V. if necessary as a corrector of hypocalcemia.

OTHER DISEASES OF UNKNOWN ETIOLOGY OTHER VISCERAL ANGIOTIDES (Diffuse Vascular Diseases)

Acute (code No 039 518)
Diffuse scleroderma (code No 114 971)
Diagnosis of polyarteritis (code No 110 0) Distal
gouty rheumatoid arthritis polyarteritis
Dermatomyositis (code No 24 100)
Pulmonary infarction (code No 460 190)

Differential

See label on page 520

Treatment

- A Spinal Muscles Norkown Supportive treatment with the use of ACTH. The effect on the may give some benefit although with a variable response the effect is quite different. The administration of corticosteroids with the lymph of

- a Low sodium diet (see page 55) and mercurial diuretics (see page 204) are of particular value in promoting diuresis and treating failure in acute rheumatic fever
 - b Digitalis is generally not as effective in acute rheumatic fever as in most cases of congestive failure and the drug may accentuate the myocardial irritability producing arrhythmias that further embarrass the heart
 - c Many cases of congestive failure are due to acute myocarditis. These cases often respond dramatically to corticotropin (ACTH) or the corticosteroids. When these agents are used for this condition maximal sodium restriction (under 200 mg daily) is imperative
- 2 Pericarditis. Treat as any acute non purulent pericarditis (see page 188). The rheumatic effusion is sterile and antibiotics are of no value. The general principles include relief of pain by opiates if necessary and removal of fluid by cardiac paracentesis if tamponade develops. If paracentesis is performed it should be preceded and followed by a short course of penicillin therapy to prevent contamination of the pericardium. ACTH and corticosteroids as well as salicylates should be continued or started as they seem to have a specific and favorable effect in aiding resorption of the fluid

Prophylaxis

The principles of prophylaxis are to avoid hemolytic streptococcal infection and to give immediate treatment with the antibiotic agents if a streptococcal infection occurs

A General Measures

- 1 Avoid contact with persons who have colds or upper respiratory infections
- 2 If possible live in a warm climate

B Prevention of Infection. The methods of prophylaxis are now indicated

- 1 Penicillin. One penicillin in doses of 200 000-250 000 units every day before breakfast. Benathine Penicillin G U.S.P. (Bicillin®) 1 200 000 units I.M. once a month. This is indicated especially for children who have had one or more acute rheumatic fever attacks and should be given throughout the school year. Adults should receive this for about 5 years after a attack of acute rheumatic fever. In any case it should be given to these individuals between September and June
- 2 Sulfonamides. If penicillin is not available give sulfadiazine 0.5-1.0 Gm. (7½-15 gr.) daily throughout the year. Patients receiving sulfonamides should have frequent blood counts; urinalysis should be performed initially and at least every 4-6 weeks thereafter. If there is any tendency towards leukopenia the drug should be stopped immediately

- C Treatment of streptococcal sepsis. Those who should be treated by one of the antibiotics. It has been shown that prompt therapy (within 24 hours) of streptococcal infection by 600 000-900 000 units of Benathine Penicillin G U.S.P. (Bicillin®) I.M. or 300 000-600 000 units procaine penicillin with aluminum monostearate in oil I.M. every third day for 3-14 days will prevent

the illness. Other patients have received temporary benefit during acute episode. A few patients seem to be only deteriorated by long-acting drugs. Suggested dosages schedules are comparable to those employed in rheumatoid arthritis (see page 315).

B. General Measures

1. Diet: High caloric, high vitamin diet.
2. Blood transfusions may be used if anemia is present. Iron salts may also be tried but are usually ineffective (page 319).
3. Protect against exposure to sunlight or other strong light (Disseminated lupus erythematosus and dermatomyositis).
4. Protect against exposure to cold (Scleroderma with Raynaud's phenomenon, see page 310).
5. Protect against secondary infections. During the acute febrile phases of disseminated lupus erythematositis and periarteritis nodosa, the administration of penicillin or other antibiotics may help to prevent secondary infection, especially to pneumonia organisms.
6. Physical therapy measures may be indicated in the management of joint and periarticular manifestations (see page 323).
7. Salicylates and other analgesics may be employed properly.
8. Proper care of skin is indicated. Steroid hormone ointments may be beneficial in scleroderma (see page 186). For care of lesions of lupus erythematosus see page 83.
9. If renal insufficiency is present treat according to general principles on page 300.

P. Prognosis

Disseminated lupus erythematosus and periarteritis nodosa usually run a rapid downhill course with a fatal outcome in 2 years. Corticosteroids or cortisone therapy appears to prolong life in a few patients. Relapses and remissions occur frequently in dermatomyositis and 50% of patients die spontaneously. Scleroderma is slowly progressive and debilitating. Acrocyanosis has a better prognosis than diffuse scleroderma.

	D	min	d	C	Supra	De	m	in	a	n	c	e	
DIFFERENTIAL DIAGNOSIS OF VISCERAL ANGIOTIDES (Modified after D. Bohr and J. Weiser)													

[illegible]

FROSTBITE (code No 0 -448)

Frostbit is i ju y of th u perfici lti su s due to fr zing it
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 3 d d g ee F eezing with de th of skin and/o d p rti s u s
 In mild c s s ther is numb n ss pri k ng and it hang With
 in r ss g e e ity the may be pare thesias and st ffa ss Du
 ung th wing t nde ne s and b ung pain a present Th skin i
 white or y ll w in c lo nd the e is stiffn s and imm bility
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T eatment

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 e lu t v ri u t e tm it m thods i e it is diffi ult to d te min
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 tent a d s ver ty f injury wh n the pat t is fir t e n

A Immediat Tr tment

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 h s d c t ted advant ges to sl w ewarming but re e t x
 p riment lw k gg sts th t mor r p d th wing t t mp
 atur sl ghtly abo e body he t may s lt in significant d
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 n t m th n 108 F (42 C) After thawing has occu d
 and th p rth s etu ned to n rm l t mpe t re xt nal
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 r d at o m l com t mp tu Avoid d e t o
 t ct with bed line bla kets a d lothi g

c A d a t p sur d e i gs and g e y handag s
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 without a ti i nfe tive g nts

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 for period of bout ew k Thi m y b s ful in
 p ev ting ond y th mbo e in s r ounding r e

g V so dli tors T e ag nts h ve not pr ved to b p
 ti ul ly valuable

B Follow p C r

1 M ld p g e ive physical the py is important a the h l
 ing p ocs oc u

2 B g e is s m y be in tit t d wh n tole t d (p g -09)

Chapter 22

DISEASES DUE TO PHYSICAL AGENTS

DISORDERS DUE TO COLD

Exposure to cold produces immediate localized vasoconstriction followed by generalized vasoconstriction. When the skin temperature falls to 25°C (77°F) tissue metabolism is slowed but the demand for oxygen remains greater than the slowed circulation can supply and the area becomes cyanotic. At 15°C (59°F) tissue metabolism is markedly decreased and the dissociation of oxyhemoglobin is reduced giving a pink well oxygenated appearance to the skin. Evidence indicates that tissue survival at this temperature is slight. Tissue death may be caused either by ischemia and thromboemboli in the smaller vessels or by actual freezing with the formation of ice in the tissues. Freezing does not occur until the skin temperature drops to -4 to -10°C (25 to 14°F) or even lower depending on coexisting factors such as wind, immobility, venous stasis, malnutrition and occlusive footwear.

Prophylaxis

1. Wear warm, dry clothing preferably several layers to afford additional insulation with windproof outer garment.
2. Keep dry when possible; remove wet clothing as quickly as possible and replace with thoroughly dried ones.
3. Avoid cramped positions, constricting clothing, and prolonged dependency of feet.
4. Exercise arms, legs, including fingers and toes periodically to maintain circulation.
5. Avoid wet and muddy ground; seek shelter from wind.
6. Maintain good nutrition and cleanliness of skin.

CHILBLAINS (code No. 0 440)

Chilblains are red, itching skin lesions usually on the extremities caused by exposure to cold without actual freezing of the tissues. They may be associated with dermatoblastoma and are aggravated by the application of warmth. In the chronic form ulcerative or hemorrhagic lesions may appear and progress to scarring, fibrosis, and atrophy.

Treatment

1. Protect affected areas from trauma and secondary infection.
2. Do not rub or massage injured tissues or apply direct heat.
3. Elevate affected part slightly and allow to warm gradually.

Prophylaxis

- 1 Avoid unnecessary exposure to heat
- 2 Maintain adequate fluid and salt intake using 0.1% saline as drinking water or salt tablets and water
- 3 Increase activity slowly until acclimated
- 4 Wear loose fitting clothing (preferably white) which is permeable to moisture
- 5 Avoid alcoholic indulgence excessive fatigue loss of sleep or intercurrent infections. Maintain good nutrition

HEAT STROKE (Sunstroke) (code No 010-453)

Heat stroke is a rare disorder due to exposure to high temperatures which is characterized by sudden loss of consciousness and by failure of the heat regulating mechanism as manifested by hyperpyrexia and cessation of sweating. There may be premonitory headache, dizziness, nausea, and visual disturbances. The skin is hot, flushed and dry and the pulse is rapid, irregular and weak. The rectal temperature may be as high as 103-112 F (42-44 C). Hydration and salt content of the body are normal.

Treatment

Aimed at reducing high temperature

A Emergency Measures

- 1 Place patient in a shady cool place and remove clothing
 - 2 Cool patient by fanning and sprinkling with water
 - 3 Immerse in cold water or use ice packs or give water enemas to reduce body temperature. Do not lower body temperature below 102 F (39 C) too rapidly
 - 4 Massage extremities to maintain circulation
 - 5 Avoid sedation since this further disturbs the heat regulating mechanism unless patient is having convulsions
 - 6 Physiological saline 1000 cc (1 qt) I.V. very slowly
- B Follow up Avoid immediate re-exposure to heat. Hyperemia likely to high temperature may remain for a considerable time

HEAT EXHAUSTION (Heat Prostration) (code No 010-445)

Heat exhaustion is due to inadequate or depletion of the physiological relation secondary to salt depletion and dehydration following sustained exposure to heat. The symptoms include weakness, dizziness, stupor, headache with or without muscular cramps. The skin is cool and pale and there is profuse perspiration, oliguria, tachycardia, with occasional mental confusion and muscular incoordination. Laboratory studies reveal hemocrit and salt depletion.

TreatmentA Emergency Measures

- 1 Place patient at rest in a cool shady place
- 2 Evaluate fluids and mass age legs
- 3 Sodium chloride 0.1% solution, by mouth or physiologically saline 1000-2000 cc (1-2 qt) I.V.

524 Heat

- C Surgery Surgical amputation should not be considered until it is clearly established that one is dealing with non viable tissue. Tissue necrosis even with black eschar may actually be quite superficial and there may be viable skin underneath which may heal well.

Prophylaxis

See General Section on page 522

IMMERSION FOOT (Trench Foot) (code No 096-44x)

Immersion foot is caused by prolonged immersion in cool or cold water or mud and is characterized by cold anesthetic extremities which become hot with intense burning and shooting pains during the hyperemic period. The affected extremities may be pale or cyanotic with diminished pulsations during vasospastic period followed by blistering swelling redness heat ecchymoses hemorrhage or gangrene and secondary complications such as lymphangitis cellulitis thrombophlebitis.

Treatment

Be instituted during stage of reactive hyperemia

A Early Treatment

- 1 Protect extremities from trauma and secondary infection. Do not rub or massage feet or legs or apply ice or heat.
- 2 Keep feet elevated to aid in removing edema fluid.
- 3 Protect pressure sites (e.g. heels) by use of pillows.
- 4 Warm extremities gradually by exposure to cool air. Do not moisten skin or immerse in water.
- 5 Bed rest early and until all ulcers have healed.
- 6 Penicillin should be used if infection develops (see page 502).

B Aftercare Treatment for Buerger's disease (see page 209)

Prophylaxis

See General Section on page 522

DISORDERS DUE TO HEAT

Exposure to heat results in prompt vasodilatation increased cutaneous circulation increased cardiac output and sweating. The resultant circulatory instability may lead to syncope in the erect position but muscular activity usually prevents this syncope. Fluid loss through sweating may amount to 3 or 4 liters per hour with heavy work at high temperatures. The salt content of sweat increases with rising temperatures ranging between 0.2 and 0.5%. Acclimatization usually results after 8 to 10 days of exposure to high temperatures but even a fully acclimatized individual may suffer a breakdown in the event of excessive fatigue. Intercurrent infection alcohol indulgence loss of sleep or failure to maintain hydration salt intake or caloric intake. Breakdown may be due to failure of the circulation (circulatory failure) or to a failure of the sweating mechanism. Cessation of sweating is a very important sign and may indicate impending stroke or collapse.

A E n r g n y M s e s (S p g 28 f t tment of sh k)

1. Plasmid transfer to once A rough estimate of plasmid
needed may be determined by one of the following formulae
Administer 100 cc of plasmid

a For e h 1% in eas in h m t rit b y 40%

rb For each 100,000 increase in RBC, boy \$ million

For the h_{27} in case in Hgb ab ve 100%

H man m lbumin app opriat do g may b sub
st tut d f r pl sm whe val ble ths d r es pos b lity
f t ansmist ng h p tit (page 26)

2 Blood t an f ns f blood lo s has oc urr d

B S g al M s r s

1 Class g and d b idem pt (A pt c tech th oughout)

a. A e that e p t nt if n ssa y

b Cleans b rn and urrounding ea g ntly with p nd
t il saline Do tr b vng o sly Eth o b ne
may b us d ior move g se ol
Remov o ly loo nd n roti t ue If n ess y to
ope blister do so sept liv

d Do not apply tartrate acid silver nitrate or sulfonamide powders to burned area

2 P s r d ring (Maintain st t a p s s)

a C r w th st rll petrol t m r mil g u st ipa

b Apply g e p d d fl fr d po ges

Band g snugly w th oll bandag d o er w th otto
p g so teril ma h n i t w t (May b t t te
Kerle ® oth r fluffy typ of b nd g f th t p)

d Apply p es u e mu l t i k i e t t l a t i b a n d g e s
R m e a f t 10 14 d y P r p e f s k i g a f t

3 Op i t t m n t i b i g u e d g a b u t h p r s s
d r s s g t h e m s t m m o l y m p l y d f o r m f t h a p y

C G I M r

1 P lin M if inf t on i s p c t e d (p g 502)

Tet s d ga g g ant tox p n (s pag 471)

3 Fl det m ntain in y tput t 1200 1500 d lly

4 S d m hl ld 8 1 Gm (2 3 d) d ly

5 Plasma blood sediment hypotension yo lat

6 High c 1 (3500 C 1) d i t with added v t min (s e pag
58)

7 The pla of t t pin (ACTH) dth ot oes t
man g m nt of b i n t l Th i ff t the ute
b do t pp t b d m t o g lly ported
Th y m y h v g t r v l d ing th ph s of h lung

ELECTRIC SHOCK (code No 010-460)

D t t m chl d g o s than lt tng r t
Alte i g t of high f q y high v lt g may b le s
dang tha l w f qu y low olt g With lt ti g ur
e t of 25 to 300 y l s low olt g (b l w 2 0) t d t p due
t c l fub llati high volt g s (1000) r spi to y
f il int m d te v lt ag (220 1000) b th El ct ic b s
a s lly h ply d m r c t ed ou d o val pal les g y r
with t c i d i n f m m t y e ti Littl happ s to them
fo ve l w k l g h i g t h n occu slowly a d n a f ly wid
a e Elect i shock m y p oduce ion of o ous s whch

4 Treat shock when present (see pag 28)

B Follow up Avoid immediate re exposure to heat

HEAT CRAMPS (code No 270 445)

Heat cramps are painful spasms of voluntary muscles of abdomen and extremities due primarily to salt depletion and following sustained exposure to heat. The skin is moist and cool and muscle twitchings may be present. The temperature is normal or only slightly increased. Laboratory studies reveal hyponatremia and low serum sodium.

Treatment

A Emergency Measures

1 Sodium chloride (tablets) 1 Gm (15 gr) every 1/2 hour with abundant of water or saline solution by mouth or 1000 cc (1 qt) physiologic saline I V. This usually relieves attack promptly.

2 Have patient rest in a cool shaded place.

3 Massage muscles gently.

B Follow up Rest for 1-3 days depending on severity.

BURNS

Burns are tissue injuries due to heat and may be graded as follows:

1st deg Erythema without blistering (code No 13 4411)

2nd deg Erythema with blistering (code No 13 4412)

3rd deg Destruction of deep tissues (code No 13 4412)

When tissue is burned plasma is lost into the burn area and fluid is sucked out of the burn. This leads to hypovolemia which continues as long as a granulating surface is present and the granulating surface heals poorly as long as the plasma hypoproteinemia. The loss of plasma results in a reduced blood volume, hemoconcentration, low renal output, decreased blood flow, oliguria, elevated N.P.N. and leukocytosis. Though anemia may occur at the time of the burn due to red blood cell destruction, it more commonly becomes apparent about the fifth day as hemoglobin and the effect of blood destruction and impaired blood formation make themselves apparent. Secondary infection is prone to occur and must be treated promptly. Death may result in an adult when 30% or more of the body surface is involved. In an infant 10% may be associated with very severe effects.

The course of a severe burn may be divided as follows:

1 Immediate shock (first 48 hours)

2 Burn shock (first 48 hours)

3 Toxemia (occurring about 3rd day)

4 Septicemia (about 3rd day)

5 Healing and restoration of function

Treatment

Take blood pressure, pulse, hemoglobin, RBC hematocrit and plasma protein at start of therapy and at regular intervals.

less often on lower abdomen or thorax and rarely over the
 t emita. The patient's psychological reaction to his illness plays
 an important contributing role.

Treatment

A Specific Measures

1. History and explanation
2. Pyridoxine Hydrochloride U.S.P. 50-100 mg ($\frac{3}{4}$ - $1\frac{1}{2}$ g)
 I.V. may be given but this is usually discontinued
3. Sedative (see page 39) and antinauseant (see page 42) drugs
 may be employed
4. Dimhydrin U.S.P. (Dormamine) 100 mg ($1\frac{1}{2}$ gr)
 20-60 mg intrab. before and repeated $1\frac{1}{2}$ - $4\frac{1}{2}$ hours
 after therapy

B General Measures

1. General supportive measures. When patient is debilitated
 symptomatic measures may be of great value.
 Electrolyte and fluid balance. Correct any deficiencies
2. Transfusion with whole blood if anemia is present

IRRADIATION SICKNESS CAUSED BY NUCLEAR RADIATION (ATOMIC BOMB)

The symptom dose relationship are with dose levels. As
 far as is known there are no permanent effects in mild cases if
 recovery occurs.

Treatment

There is no known perfect treatment. General supportive
 measures with complete bed rest and quiet nutrition as for
 febrile patients when indicated and blood transfusion when hypotension
 occurs are all that can be offered at present.

Prophylaxis

The most important danger of an atomic explosion is the thermal
 fire. This can be minimized somewhat by fleeing to the ground
 or going to the exposed part of the body or seeking shelter behind a
 building wall or trench. For the 10 seconds immediately after the
 bomb explosion. The following are measures advocated by the
 Atomic Energy Commission to minimize exposure to radiation.

- A. Clothing will provide a measure of protection to the skin.
 Any contaminated clothes should be disposed of as soon as
 possible and before entering a non-contaminated area.

B. Clean All Exposed Skin

1. Vigorously rubbing with soap and water rapidly syn-
 thetically detergent pyrolytic particulate matter to the hair
 and skin fold and as about body orifices will give
 first degree contamination. Do not abrade skin and
 do not use contaminated water.
2. If soap and water are not effective isotonic saline (pH 7.0)
 or a mixture of boric acid and starch will be useful.
 Sodium bicarbonate solution is also recommended especially
 for mucous membranes.
3. In an emergency wipe skin with any non-contaminated paper
 or sand and sweep grass leaves etc.

may be momentary or prolonged. With recovery there may be muscular pain, fatigue, headache, and nervous irritability. The physical signs vary according to the action of the current. With ventricular fibrillation no heart sounds or pulse can be found and patient is unconscious. The respirations continue for a few minutes becoming exaggerated as asphyxia occurs and then ceasing as death intervenes. With respiratory failure respirations are absent and the patient is unconscious; the pulse can be felt, but there is a marked fall in blood pressure and the skin is cold and cyanotic.

Treatment

A. Emergency Measures

1. Free from current at once. This may be done in many ways but rescuer must protect himself in the process. Turn off power. sever the wire with a dry wooden handled axe, make proper ground to divert current, or drag victim carefully away by means of dry clothing or leather belt.
2. Artificial respiration must be started immediately (see page 150) if victim has slow or absent breathing and continued until spontaneous breathing returns or rigor mortis sets in.
3. Precordial compression for ventricular fibrillation or arrest. Artificial respiration will not restore normal heart beat and other measures may not either. If possible incision of the chest and manual pumping of the heart may be employed as a last resort. Electric defibrillators may be employed if by chance available.
4. Treat shock promptly (see page 28).
5. Positive pressure oxygen with carbon dioxide may be used when available or oxygen and carbon dioxide by mask combined with artificial respiration.

B. Hospital Measures

1. Hospitalize patient when required and observe for sudden cardiac dilatation or secondary hemorrhage.
2. Lumbar puncture if signs of increased pressure are noted.
3. Treat conservatively. The direction and extent of tissue injury may not be apparent for weeks. Infection is usually not a problem early. Patience and delay are important in treatment, allowing anation tissue to be well established before attempting any surgery. Hemorrhage may occur later and may be severe.

IRRADIATION SICKNESS

Irradiation sickness is the term applied to the syndrome developing during or after the course of therapeutic x-ray administration or after exposure to ionizing radiation (e.g., x-rays, neutrons, gamma rays, alpha or beta particles).

IRRADIATION SICKNESS ASSOCIATED WITH IRRADIATION THERAPY

Anorexia, vomiting, weakness, exhaustion, lassitude and in some cases prostration may occur singly or in combination and may be of varying severity. The symptom complex is most likely to occur when x-ray therapy is given in a pro-

Collect and s v washings in clean c tainers fo toxic o
l g l e amination when dicated In f e sic ca es s al
w th ealing wax and pla e in a locked r frigerato deliv r
to toxic logi tpe sonally a d g t a sign d rece pt If re
fr ge t ni la king pre erv the sp cimen w th equal
quant ti of 95% alcoh l do not e formalin as this int
fer s with toxicologic xami tion

- 4 Gastr la ag fluids (1) Wa m tap wat or 1% saline
- (2) Thin sta b pa te ol bl (3) Sodium bic rbo te 1%
- (4) Pota alum permang ate 1:2000 (1 Gm in 2000
wat) (5) Sod: m th o ulf te 1% (6) Hydrog p ox de
1 or 2%

C C tha is May b ff tiv in ret ding ab ption

N t al t on (I ctivation) of P s P or to Intest al Abs ption

Ex ption s of t g a d or alkals alw ys f ll w th
g at cl v g

A N ut al ation of Ac d d Alk l s See sp f c pos

B P cipitation f Ch mi l S e pec f poiso

C l t t by D mul nt D mul nts p ipitat metals and
l h lp dimin sh ab ption of many poisons Th s bland
g nt a eals oothi g t inflam dm o memb anes U
3 o 4 aw gg whit be t in 500 c milk o water kimmad
milk thun flour at rch solut n (bo l d if pos ble)

S y p t o m s & Symptom tic M es

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b rvati n in o de to ticip te the imm di te and d lay d com
pli at o s f the p isoning Sui dal pat t may d p iate r
eulla c and should b pla d d the a of a p ychiat it

A C culat y Fail e

- 1 Sh k (s p g 27) Th prin ipal m includ re
cumb ntpo t n wa mth dministration of tum lants and
p r t al fluid to n effectiv bl od v l m
- 2 Ca d l f ul e (p ge 181) The p in ipal m ur in
l d oxyg n d g t h m r l d ur tics and re ely
tr ardi p n ph me
- 3 Pulm n y dem The p p l m s in l d oxyg n
p i lly by po ti e p ssur (p g 147) o t eatm nt
fo di failu f t x i t vo d p nt al alin o
oth p t l fluid (x ept pl sm)

B R p tory Abno m l t

- 1 R p tory b t tion Co t (by opharyng l irway
i tr t h l int bation o t h otomy
- 2 R p t y dep i Pla in op n ai Admi t
a tifi al e pur t p R us it to o oth r mean of
tomat ntill tu sh uld b mpl yed soo a po ple
St mulants (anal pti dr g) a of d t: abl val e n
w th C N S d p s ant drug
a W m st ong bla k off o ally o ctally
b W m stro g t a o ally
A omati pl it of ammonu 2 4 (1/2 l d) in l c p
f w t
- d Eph dri Mat 50-120 mg (3/4 2 g) orally o b ut
N k th mid inj tio (C mi) 0 25 1 25 Gm (3 3/4
18 3/4 gr) I V

DISEASES DUE TO TOXINS

PRINCIPLES OF TREATMENT OF ACUTE POISONING

In the emergency treatment of a *any poisoning* in which the toxin has been taken by mouth the following general procedures should be carried out: (1) Remove poison by emesis, lavage, catharsis or diuretics as soon as possible. (2) Inactivate poison with specific or general antidote. Follow with lavage. (3) Combat shock, collapse and specific manifestations as they arise. (4) Protect mucous membranes with demulcents.

Removal of Poison

Do not use stomach tubes or emetics in poisonings due to strong acids or alkalis or other corrosive agents they may cause gastric perforation.

A. Emesis. This is the quickest way to evacuate gastric contents.

1. Indications. For removal of excess poison in cooperative patients or for convenience when a stomach tube is unavailable or patient is unable to take stomach tube.
2. Contraindications. (1) Drowsy or unconscious patients (danger of aspiration of stomach contents). (2) Ingestion of corrosive poisons, kerosene or convulsants.
3. Technique. Introduce finger, feather or other object into throat or give one of the following and follow with copious quantities of warm water. (Emesis should be continued until gastric contents are clear.)

Apomorphine hydrochloride 5 mg (1/10 gr) subcutaneous will often quiet the patient and will usually induce vomiting.

- b. Powdered mustard 1-3 tsp in a glass of lukewarm water is an unpleasant and unpleasant emetic but is often safe and has the advantage of being generally available.
- c. Sodium chloride 1 Tbsp in a glass of lukewarm water is not very effective but is readily available.
- d. Stomach pumps 250-500 cc.

B. Gastric Aspiration and Lavage

1. Indications. (1) Removal of excess of noncorrosive poison which may later be absorbed from the gastrointestinal tract. (2) Removal of CNS depressant poison when vomiting does not occur (omitting cathartics). (3) For collection on a deamination of gastric contents for identification of poisons. (4) For convenient administration of antidote.
2. Contraindications. (1) Extensive corrosion of tissues by poison. (2) Staggering delirious stupor or coma. Patients be cause of danger of aspiration pneumonia.
3. Technique. Gently insert a lubricated soft but non collapsible stomach tube through the nostril into the stomach. Lavage copiously but do not distend the stomach. Under no circumstances it is better to lavage with a small quantity of fluid at frequent intervals. Always remove excess of lavage solution.

sw allowed a c rosi e poison (symptom sev e pain burn
ings sation i mo tha d throat vom tng) CALL PHY
SICIAN IMMEDIATELY Acid d acid like co rosi v in
lude sodium a id sulfate (toilet bowl cl a ers) acetic acid
(gl a al) sulfu i acid nit ic acid oxalic cid hydrofl o ic
a id (rust r mo s) iodi e a d s l e r n t ate (styptic p n
cil) Alk li rro i es includ od m hyd ox de ly (drain
cl s) sod um carbonate (w hing oda) mmonia wate
nd od um hypochlo ite (household ble h)

If the p tie t can swallow aft i g at ng a corrosiv poi
son th foll w g s b tanc s (and amount) may b giv

For ids Milk w ter o milk of magn sia (1 Tbsp /
1 c p wat)

b F ralk lls Milk water y fruit j ce o i eg r

For p tie t 1 5 ye old give 1 to 2 p fo p tie t
5 y a d olde p to 1 q rt

- 2 I d e vom tng when on cor o ve sub t ces have b en
swall w d Gv milk o w t r (fo patie t 1 5 y s old
1 to 2 c p fo p t ent ver 5 ye rs up to 1 q art) I d c
v m tng by pla g f ng or the bl nt e d of a spoon t
th b ck of th pat ent throat or by the u fa m tic
(2 Tbsp s It i a gl s of warm wat r) Whe tching d
vomit g begun place p t t fa d wn with h ad low than
h ps This pre e t vom tu from t r ng th l g d
c i g f ther dam g

- B Inhal d Pois ns C rry pat ent (d not l t him walk) t f e sh
al mmedi t ly Open all doo and windows Loosen all
tight lothing Apply a tif ial r p r tion if b thing h
stopped or is ir egul r P v t chilli g (wrap pat nt in
blank ts) Keep p ti nt as qui t as pos ible If p ti nt is con
vulsing k p him in bed in a semi d rk room avoid jar ing o
n i e D not giv alcoh l in any fo m

- C Sk C t m n t D ch skin with wat r (how r hos
f t) Apply t e m f wate on skin while remo ing cloth
ig Clean e skin th ro ghly with wat p dity w hing is
m t important in r ducing xt nt of i j ry

- D Eye Co t m ti H ld y i d pen w hey with g tl
t am of nn g w te immediately D lay of a f w cond
g tly c es t nt of inj y Co t w shing til phy
sicia a Do not use ch mi als they m y i as
exte t of njury

- E l i f e t d Poi n (5 o p ion and ake bit) Mak patient lie
dow as soon possibl Do ot giv alc hol in any f m
Apply tourniq t bov inj ti nait (i b twe th bit
and th h a t) Th pul in v ls b low th tou niqu t sh ld
ot di appe hould th to r niqu t prod c a throbbing
ns ti n T urniq t hould be loos ned f r l minut e y
15 minut s Apply ice pa k to the s t of th bit Carry p
ti nt to physician o ho p t l Do n t let h fm walk

- F Ch mi al B rn W sh with l g quantities of runni g water
(pt in th c f b ns aused by pho ph) Cover
imm di t ly with loo ly ppli d l an loth Avoid u e of
ointment gr as s powde s and oth r dr gs in f st aid treat
ment of b rn T t h ck by k eping pati nt fl t k pung
him w m nd a ring him until arri al of phys ian

- f Amphetamine sulfate 5 40 mg ($\frac{1}{12}$ $\frac{2}{3}$ gr) orally or I V
 g Methamphetamine hydrochloride 2 5 15 mg ($\frac{1}{24}$ $\frac{1}{4}$ gr) orally or I V [For use of pentylenetetrazol (Metrazol®) and picrotoxin see page 537]

3 Hypostatic pneumonia [see page 122] The principal measures include antibiotics and intratracheal aspiration p r n

C Central Nervous System Involvement

1 C N S excitement Use hypnotic or anticonvulsant drugs (see also pages 354 355) e g

- a Amobarbital Sodium (Amytal®) 250 500 mg ($3\frac{3}{4}$ $7\frac{1}{2}$ g) as f sh 10% solution I M or I V
 b Paraldehyd (1) Oral 5 15 cc (1 4 dr) in cracked ice with milk juice or whisky (2) Rectal 5 30 cc (1 8 dr) in equal quantity of vegetable or mineral oil (3) I M 5 10 cc i to buttock
 c Calcium gluconate 10% 10 20 cc ($2\frac{1}{2}$ 5 dr) I V

2 C N S depression Use stimulant drugs (see above)

D Dehydration Use oral or parenteral fluids as tolerated and indicated (see page 18)

E Pain See analgesic and narcotic drugs on pages 32 and 33

A M.A. RECOMMENDATIONS ON FIRST AID MEASURES FOR POISONING*

[Since prevention and helpful first aid measures are of great importance the following is provided for the physician's use in public education]

The aim of first aid measures is to help prevent absorption of the poison. SPEED is essential. First aid measures must be started at once. If possible one person should begin treatment while another calls a physician. When this is not possible the nature of the poison will determine whether to call a physician first or begin first aid measures and then notify a physician. Save the poison container and material itself if any remains. If the poison is not known save a sample of the vomitus.

Measures To Be Taken Before Arrival of Physician

A Swallowing Poisons Many products sold in and around the home although not labeled poison may be dangerous if taken internally. For example some medications which are beneficial when used correctly may endanger life if administered properly or in excessive amounts.

In all cases except those indicated below REMOVE POISON FROM PATIENT'S STOMACH IMMEDIATELY by inducing vomiting. This is the essence of the treatment and is often a life saving procedure. Prevent chilling by wrapping patient in blankets if necessary. Do not give alcohol in any form.

- 1 DO NOT induce vomiting if patient is in comatose unconscious or having convulsions if he has swallowed petroleum products (i.e. kerosene gasoline lighter fluid) or if he has

*Recommendation of the Committee on Toxicology American Medical Association. Bernard E. Conley Ph.D. Secretary. Modified and reproduced with permission from 887 7

ALCOHOL METHYL (code No 010 331)

Methyl alcohol is a common and instant and CNS depressant which has an affinity for the optic nerve. It is slowly excreted from the body and is metabolized giving formaldehyde and formic acids as products which produce as follows. The M.L.D. by ingestion is 30-60 cc (1-2 oz). The patient experiences headache, epigastric pain, dyspnea, and vomiting and may suffer loss of vision. Examination reveals hyperemic conjunctivae, extremities depressed, delirium, coma and convulsions.

Treatment

1. Give stomach wash with 1 or 2% sodium bicarbonate.
2. Check serum CO_2 .
3. Combat acidosis. Give sufficient sodium bicarbonate orally or alkalinize by intravenous maintenance alkali such as serum CO_2 content of over 20 mEq/L.
4. Keep patient in shaded room; give supportive measures as required.
5. Administer Ethyl alcohol 100 proof 50% 3-20 cc very 2-4 hours for 3-5 days.

ALKALIS (code No 010 32)

The strong alkalis are common ingredients of household cleaning compounds and many industrial cleaning agents. They irritate the effect of the known domestic members but in instances of severe poisoning such as industrial and oil poisoning. The M.L.D. for the alkali powder (NaOH and KOH) is 15 Gm ($\frac{1}{2}$ oz) if at once mixed with water 4 (1 d). Recovery may follow if much large doses how ever. The symptoms of burning pain, the appearance of the skin, the use of vomiting and difficulty in swallowing and burning. Physiological examination reveals dehydration and edema of skin and mucous membranes in and about the mouth, bloody vomit and stool, dyspnea and pulmonary edema.

Treatment (Administer antidotes if possible as a possibility)

1. Dilute in gastric juice; neutral alkali 120 to 240 (4-8 oz) of 5% hydrochloric acid may be used.
2. Give fluid orally (helps excretion by formation of soap) white flour egg white mixed with water 1-2 Tbsp given 300 cc (1 pt) if water.
3. Withhold food until vomiting has ceased.
4. Supportive measures as necessary.
5. Withhold food with dilute vinegar or fruit juice.
6. Wash eyes with water.

ARSENIC

(Artenide No 010-3114) (Chronic code No 011 3114)

Arsenic is found in industrial chemicals, pesticides and as a natural mineral.

A variety of the following may occur: nausea and vomiting, abdominal pain, diarrhea, a marked thirst, burning sensation and difficulty in swallowing. Cyanosis may develop, abdominal symptoms common.

Measures to Prevent Poisoning Accidents

- (1) Keep all drugs, poisons & substances and household chemicals out of the reach of children
- (2) Do not store noxious products on food shelves
- (3) Keep all poisonous substances in their original containers do not transfer to unlabeled containers
- (4) When medicines are discarded discard them. Do not throw them where they might be reached by children or pets
- (5) When giving flavored or brightly colored medicine to children always refer to it as medicine never as candy
- (6) Do not take or give medicine in the dark
- (7) READ LABELS before using chemical product

TREATMENT OF SPECIFIC POISONINGS (ALPHABETICAL ORDER)

ACIDS CORROSIVE (code No. 010 32)

The strong mineral acids exert primarily local corrosive effect on the skin or mucous membranes. In severe burns result to yolk pus may result. The M.L.D. is 4 cc (1 dr.) of concentrated acid. Symptoms include severe pain in throat and upper gastrointestinal tract, marked thirst, bloody vomitus, difficulty in swallowing, breathing and peking discoloration and destruction of skin and mucous membranes in a day or two, mouth collapse and shock.

Treatment (Avoid emetic or lavage if possible, if it is a possibility)

- 1 Dilute acid immediately with copious amounts of water, milk or egg whites. Do not waste time looking for specific chemical antidotes.
- 2 Limestone, magnesia or aluminum hydroxide orally to neutralize acid. Avoid carbonate or bicarbonate internally if possible since they form gas and distention of already weakened stomach wall.
- 3 Egg white beaten with 500 cc milk or water as demulcent.
- 4 External burns. Wash with water and remove debris.
- 5 Eye. Wash well with 1% sodium bicarbonate solution.
- 6 General supportive care as detailed (page 531).

ALCOHOL ETHYL

Ethyl alcohol is a mucous membrane irritant and a CNS depressant. The M.L.D. is 100-200 (3-7 oz.) if pure alcohol when ingested at one time.

Treatment (Avoid sedatives and narcotics)

A. Acute Intoxication (code N. 010 33)

- 1 Careful examination especially for hypothermia
- 2 Large quantities of warm water containing 4 Gm (1 tsp) sodium bicarbonate given symptomatically 6 mg (1/10 gr) if alcohol has recently been ingested.
- 3 Stimulant. Strong black coffee orally or rectally. Naloxone injection (Coramene®) 25 mg 1-3 ml IV.
- 4 Resuscitation for comatose patient if dead immediately. Cause of death is usually respiratory failure.

B. Delirium Tremens (code N. 003 332)

- 2 Gastric lavag with 1:2000 pta (umpe manganate B certain t remove all permangan te in final pir tion Thi s of doubtful value if perform d mo e than 6 hou s after in gestic n and may be dang ous CAUTION D nge of asp r ati n pne monia is great in stupo ou or comato e patients
- 3 P gation is of no value
- 4 Indwelling catheter Sav all urine fo toxi ologic st dies
- 5 Antibiot dr g P nicillin 300 000 to 500 000 units I M daily to less n dang of pne m ni
- 6 P teral fluid If diac failure i absent and renal fun tion is adequate giv 1 L physiologi al line and i 2 L 5% d xtro e I V in w i r d ily to maintai rine output (1 i 5 L /24 h u s) U less fluid loss has b n excessi don t gi e mo th 2 3 L f fluid du ing the first 24 h rs In e ent of ci c lat ry collap use plasm
- 7 C ntral nerv us sy tem stimulants (anal ptics or convulsant d g) These e n t true ba bit te art dote but ar utili ed to maintain patient s r p tions and reflex s Th ir pl c in th apy i uncertain They do ot ho ten th d ration f ffect of the drug and fter th i se th d p e sion m y be ev mo ev r They ar ften dang ou dr gs a d unless used carefully m y j opardiz the patient s hanc for r very S pe ority of th va o anal pti agents is o tro r i l so alar m ntio d CAUTION Anal pti s should n t be cons d to substit f di ect phyologi al t e tment of dep essed r spi tion a d c cul tion in b bit rat poi ning
 Pic toxin Inj tio U S P (0 3% o 3 mg /c)
 Administer 2 3 I V (o I M) at onc a d follow w th 3 c v y 20 30 ma t until t rn of fle small twit hing or body m veme t (ot o vuls ons) G m intenance d s ne e s ry to keep patient at thi lev l
- b P tylen t t az i Inj ecton U S P (M tra ol[®]) (10% o 100 mg /c) (Aft r M phy et l) Give 3 5 cc I V at o 15 c I V in 15 minute if p ti t is ar flexic 20 I V ev ry 30 min tes until fl turn and 2 5 I M p r n until full con cious s is st ed Oth prepa tions Amphet maine eph drin m th m phetamine and stry hnine ha e be n eds es fully alo e or n ombination but th i s pe ority o the above tre tme t m thods h not b en p ov d M tha tharumide (Megtimid[®]) (B B methyl thylgi t imid) has bee athe e thusiastically r ported s a ffective n w ps bit ate a tag nist in the t atm t of a t ba bi tur t poi ning E pe i ne in th co ntry s gg sts that the g tpo sses o specif nal ptic prope ties but th is little viden of true b bitur te tago ism
- 8 The artifi l kidney or pe itoneal dialy is i indicat d in v s when a ilable

BELLADONNA DERIVATIVES

(At pin code No 010 352) (Scopolamine cod No 010-379)

Th belladonna alkaloids a e parasymp thetic d pr ssants with variabl C N S ff ts M L D is 2 8 mg (1/30-1/10 gr) of at o line the usual lethal dos is n r 100 mg (1 1/2 g)

Treatment

- 1 Emetic or abundant gastric lavage with warm water
- 2 Follow with demulcent drink
- 3 Symptomatic relief of diarrhea (e.g. codeine)
- 4 Dimercaprol Injection U.S.P. (BAL®) 10% solution in oil
Give 1 M (Eagle J Ven Dis Inform 27:114 1946)
 - a Severe poisoning 3 mg /Kg / dose (1.5 cc /80 Kg)
 - 1st day 1 inject on eve y 4 hours day and night
 - 2nd day 1 injection every 4 hours day and night
 - 3rd day 1 injection every 6 hours for 4 doses
 - 4th day on 1 injection b i d for 10 days or until recovery is complete
 - b Mild poisoning 2.5 mg /Kg /dose (1.5 cc /80 Kg)
 - 1st day 1 injection every 4 hours for 4 doses
 - 2nd day 1 injection every 4 hours for 4 doses
 - 3rd day 1 injection b i d
 - 4th day on 1 injection once or twice a day for 10 days or until recovery is complete
 - c Toxic reactions to BAL® These appear to be transient and over in 30 minutes. They include nausea vomiting headache generalized aches and pains and burning sensation about the head and face. Barbiturates have been recommended for severe side effects.

BARBITURATES (code No. 010 3371)

Barbiturate are used for daytime hypnotic or anticonvulsant purposes. The barbiturates are one of the most common means of both suicidal and accidental poisoning.

Obtain data on dosage and time of ingestion from patient relatives, friends or attending physician when possible.

A Mild Symptoms Drowsiness mental confusion headache there may be euphoria or irritability.

B Moderate or Marked Symptoms Delirium stupor shallow and slow respirations circulatory collapse cold clammy skin cyanosis pulmonary edema dilated and non-reacting pupils hyporeflexia areflexia coma and finally death.

Treatment

- A Mild Symptoms** Symptomatic and supportive nursing care. Stimulants should be limited to caffeine. Keep patient under observation until danger has passed. Place suicidal patients under psychiatric care.
- B Moderate or Marked Symptoms** Most of the patients will survive even days of unconsciousness if the airway is kept open (usually requires tracheotomy) and if artificial respiration is maintained (usually with a tank respirator). The patient should be hospitalized and anticholinergic resuscitated (see page 27). Record the following at 15- to 30 minute intervals until the danger has passed: temperature pulse respiration and BP mental status or state of consciousness skin color (cyanosis or pallor) lung bases (pulmonary edema) reflexes (corneal pupillary gag patellar) and sensation (response to pain).
 - 1 Airway Aspirate mucus pull tongue forward and insert oropharyngeal airway. Insert tracheal or tracheostomy intubation may be advisable.

CARBON TETRACHLORIDE (code No 010 33411)

This agent is a very common solvent and disinfectant in industry and the home. It is a local irritant, cardiocirculatory and general CNS depressant and a general protoplasmic poison which has a marked effect on the liver and kidneys. It enters the body by ingestion and inhalation. The M.L.D. is 4 cc (i.d.) when ingested; the M.L.D. by inhalation is unknown. The symptoms include headache, hemorrhage, nausea, vomiting, diarrhea, abdominal pain, weakness, vital disturbances, neuritis and intoxication. Early signs are tenderness, jaundice, oliguria and edema; later epistaxis and hemorrhage.

TreatmentA Acute Poisoning

- 1 Remove from exposure; keep warm and warm.
- 2 Lavage copiously with 1:2000 potassium permanganate.
- 3 Sodium bicarbonate 30 Gm (1 oz) in water at once.
- 4 Treat as potent liver hepatitis (see page 272). Observe for oliguria. If it becomes manifest, treat as a ureteric filtrate (see page 303).
- 5 Initiate supportive therapy.

B Chronic Poisoning (Acute alcohol)

- 1 Remove from exposure.
- 2 Carefully watch heart, liver and kidney function.
- 3 Treat as potent liver (see page 281).
- 4 Symptomatic and supportive measures.

CYANIDES (code No 010 353)

Hydrocyanic acid and the cyanides are death by inactivation of the primary respiratory utilization of oxygen by the tissues. The M.L.D. is 2 (i.d.) by ingestion or inhalation. The initial period of giddiness, loss of muscular power and stupor is accompanied by panting, apnoea and profound hypoxia. The odor of bitter almonds is on the breath.

Treatment: Work rapidly for death occurs quickly

- A If i.h.i.d. (1) Place in open air; keep warm; stop motion.
- (2) Maintain artificial respiration manually until arrival of resuscitator.
- (3) Amyl nitrite (place pea in the mouth) by inhalation for 15-30 seconds every 2-5 minutes.
- (4) Sodium nitrite 10-15 cc of 3% solution or 50 cc of 1% solution I.V. at 2-4 minutes for injection.
- (5) Followed immediately with 50 cc of 25% solution of sodium thiosulfate if tolerated I.V.
- (6) Repeat N₂ and N₂S if symptoms recur.
- B If i.g.i.d. Give amyl nitrite as above; lavage stomach copiously with 3% hydrogen peroxide solution; 10% sodium thiosulfate solution; 1:2000 potassium permanganate solution.
- C Supportive therapy.

CHLOROPHENOTHALENE (DDT) (code No 010 3)

DDT is a CNS stimulant which causes poisoning by ingestion, inhalation or contact. The M.L.D. is probably about 20 cc (i.d.) but few fatalities have been reported. When

The patient complains of dryness of mouth thirst difficulty in swallowing and blurring of vision. The physical signs include dilated pupils flushed skin tachycardia fever delirium delusions paralysis and stupor.

Treatment (Avoid opiates)

- 1 Tincture of iodine 4 cc (1 dr) in 1000 cc (1 qt) of water
- 2 Universal Antidote charcoal in water (See back cover)
- 3 Lavage well with 1:2000 potassium permanganate solution
- 4 Sodium sulfate 30 Gm (1 oz) in water
- 5 Pentobarbital sodium 0.1 Gm (1½ gr) for excitement
- 6 Institute supportive measures

BROMIDES

(Acute code No 010 3217) (Chronic code No 011 3217)

Bromides are CNS depressants frequently found in medicinal preparations. Acute poisoning is rare. The symptoms include ataxia constipation drowsiness apathy and hallucination. The physical examination reveals decreased reflexes conjunctivitis foul breath frothy tongue sordes unequal and irregular pupils ataxia abnormal reflexes (often bilateral) toxic psychosis delirium and coma.

Treatment

- 1 In acute poisoning lavage copiously with saline to remove unabsorbed bromides and later to remove those absorbed into the stomach. Follow with magnesium sulfate 30 Gm (1 oz) in water for laxation.
- 2 Sodium chloride 6-12 Gm (90-180 gr) daily in addition to regular dietary intake 1000 cc saline IV bid or the same orally 0.1-2 Gm (15-30 gr) every 4 hours orally. Treat until blood bromide level is below 50 mg/100 cc.
- 3 Force fluids to 4000 cc daily.
- 4 Use continuous warm baths (95-96°F) or sedative cold packs as necessary.

CARBON MONOXIDE (code No 010 352)

This gas is responsible for many deaths and numerous deaths result from the use of unvented gas or coal burning heaters. It is also used for suicidal purposes. It combines with hemoglobin to form relatively stable compound which conditionally causes tissue anoxia. Manifestations are headache faintness giddiness tinnitus vomiting cherry red skin vertigo loss of memory fainting collapse paralysis and unconsciousness.

Laboratory Data When boiled or when shaken with 1 to 2 volumes of sodium hydroxide blood remains red while normal blood becomes black or brown black.

Treatment

- 1 Remove patient to fresh air keep warm loosely clothing and maintain rest.
- 2 Artificial respiration or resuscitation 100% oxygen per
- 3 Give 50 cc of 50% glucose IV for cerebral edema.
- 4 Institute supportive measures.

Treatment

- 1 Starch flour raw egg white or 1% sodium thiosulfate in water
- 2 Follow with emetic or remove by lavage with 1% sodium thiosulfate solution. Repeat until evidence of iodine has disappeared from gastric contents
- 3 Then give demulcents e.g. milk or barley water
- 4 Symptomatic and supportive measures for systemic reaction e.g. stimulants or anticonvulsants

LEAD (code No 010-3112)

Lead may poison by ingestion or inhalation of its dust or fumes. It has local irritating action and a generalised toxic effect. The M.L.D. is 10 Gm (150 g) of lead acetate. Poisoning is manifested by metallic taste, dry throat, thirst, abdominal colic, vomiting, diarrhoea, constipation, headache, leg cramps, black stools (lead sulfide), oliguria, stupor, convulsions, pallor and coma. In the chronic form there is a variable element of the C.N.S. blood-forming organs and gastrointestinal tract.

Treatment**A Acute Poisoning. Do not use BAL®**

- 1 Lavage with dilute magnesium sulfate or sodium sulfate solution to precipitate insoluble lead sulfate
- 2 Treat symptomatically. Avoid narcotics. Treat colic with colloidal atropine and sedatives
- 3 Edathamil Calcium Diethylenetriamine N.N.D. (EDTA Versenate®) forms a soluble non-toxic soluble complex that is excreted in the urine. It has been used successfully in the treatment of lead poisoning. Give orally I.V. (2% solution) or intravenously I.M. (20% solution containing 0.5% procaine) a total dose ranging of 50-75 mg/Kg body weight per 24 hours for a course of 5-7 days. The drug is nephrotoxic and should not generally be given to children. Giving 5 Gm per 24 hours rapidly and carefully volumes of dilution containing 1 litre may give a total daily dose of 100 ml. A analgesic and sedative for encephalopathy especially in children. Renal function and fluid and electrolyte equilibrium must be considered on an individual basis.

B Chronic Poisoning

- 1 Remove patient from exposure
- 2 Adequate diet with vitamins
- 3 Courses of EDTA may be employed especially when haematological complications have occurred (see above)
- 4 If phyllic EDTA should not be used as a substitute for properly prepared and protected emulsions (see above)

MERCURY (code No 010-3111)

Mercury poisoning occurs by ingestion or inhalation. It is a general protoplasmic poison. The M.L.D. is about 70 mg (1+ gr) of mercuric bichloride. The manifestation in the metallic taste, lividation, throat burning, sensation in throat, diarrhoea and odour of oral tissues, abdominal pain, vomiting, bloody diarrhoea and shock. In the chronic form there is weakness, ataxia, intention tremor, irritability, depression and muscular cramps.

poisoning occurs from the material in solution the actual poisoning is usually due to the organic solvent and not DDT. The manifestations are tired and aching limbs, nervous irritability, mental sluggishness, muscle twitchings, convulsions, and coma.

Treatment: (Avoid epinephrine; may cause ventricular fibrillation)

- 1 Universal antidote at once if available (see back cover)
- 2 Lavage with large quantities of warm water
- 3 Sodium sulfate 30 Gm (1 oz) in water
- 4 Pentobarbital sodium 0.1 Gm ($1\frac{1}{2}$ gr) orally. Give 0.25 to 0.5 Gm Amobarbital Sodium N.F. (Amytal®) as fresh 10% solution slowly I.V. or I.M. for convulsions
- 5 Calcium gluconate 10% 10 cc I.V. for convulsions
- 6 Supportive measures as necessary
- 7 High CHO high protein diet with added vit. B to protect liver

FLUORIDE POISONING (code No. 010 3215)

Fluorides are found in agricultural poisons and insect powders and are used in the aluminum industry. Clinical features include nausea, vomiting, colicky abdominal pain, diarrhea, cyanosis, C.N.S. excitement, and convulsions.

Treatment:

- 1 Lime water or other soluble calcium salts orally in large quantities
- 2 Give emetic or use copious gastric lavage with lime water
- 3 Egg whites beaten with 500 cc (1 pt) milk or water
- 4 Stimulants (see page 531)
- 5 Calcium gluconate 10% 10 cc I.V. repeat if tetany occurs
- 6 Give artificial respiration and combat shock

GASOLINE AND RELATED COMPOUNDS (code No. 010 33x)

Gasoline poisoning may result from inhalation or ingestion, but more severe symptoms result from inhalation because the C.N.S. is more quickly reached by this route. Acute manifestations are vomiting, vertigo, muscular incoordination, weak and irregular pulse, twitchings, and convulsions. In the chronic form there is also headache, drowsiness, dim vision, cold and numb hands, weakness, loss of memory, loss of weight, tachycardia, mental dullness, or confusion, so-called metamorphoses, and secondary anemia.

Treatment:

- 1 Remove to fresh air
- 2 Lavage with salad oil and/or large amounts of warm saline
- 3 Sodium sulfate 30 Gm (1 oz) in water followed by mineral oil 120 cc (4 oz)
- 4 Watch closely for 3-4 days for severe symptoms or collapse

IODINE (code No. 010 3218)

Clinical features include characteristic stain of mouth and odor of breath, yellow or bluish orifice, pain and burning in pharynx and esophagus, marked thirst, diarrhea (stools may be bloody), weakness, dizziness, syncope, or convulsions.

Mushroom Poisoning

	Aman ta m c i (fly aga)	Aman ta ph lloid s (dest oying ang l)
Pharm o- lg a tion	P symph th ti stimu- tion by alkaloid musca ine	Dr ito ti on i most all c lls e peci lly liv h a t kid y
U et symptoms	o dd n l 2 h urs Cnf o ex tem t thi t n ea a d vomit ing d rh wh zing s li at o al wp l tremo s weak s ollap e nd e d ath	Del yed 12 24 h ur Cnfus d p ess h dach onvul ions com na a vomiting bloody vom t a d tool p i ful larg ment f l j u dce ol g ri p lmon ry dem
R t i of t atment	1 R mo l of Gl t t by em is and lav ge fol low d by ath reis 2 C te tio of m a alk l id by s of t pi ulf t 1 2 mg (1/60 1/30 gr) ub t at on d p t q 30 min p r 3 Sed t on w th b b t at f e t m t 4 F e flu ds by o al nd p re t al te 5 T e t f h k	1 R n l of Gl o tent by me and la ag f l low d by athar 2 Ant d te Tr t non sp lli p rasymph th tic a t m ff t w th at p n l f t 1 2 mg (1/60 1/30 gr) b t at n a d q 30 mi p r n 3 R lev p n w th a ot q r n 4 Prot t li w th 4 5 l te of 5% de t e v y 24 ho r if al fun t on d qu te 5 T at f h k

OXALIC ACID (Cont d f m p vio s pag)

T ime t

- 1 G e at o l ml taste oth r lt 30 Gm (1 oz) in
wat agl s flme w t r o lag m unts f milk to
p pit te s lubl alc mo lat
- 2 L g w th l 2000 pot mpe m g nate s lot
- 3 White of ggs b t m l k a demul nt
- 4 C l i m gl te o l t te 10 c of 10% l tio I V
a d l m alt 1 2 Gm (15 30 gr) or lly q d
- 5 Inat t t s pp t v m s es

PESTICIDES

Common P st d d Th T im t

A M t l

- 1 A l [Cal i m a nate le d a e te c pp a ct
t (P is g en)] Dm p ol U S P (BAL®)
th py (p g 536)
- 2 Le d alts [L ad r t] Ed thami Cal i m D sodium
N N D (V e te®) (S e p g 541) DO NOT USE BAL®
- 3 Coppe lts [Coppe lfat (bl vit iol ed in
B r dea mixt)] Symptomatic and s pporti Sodi m

Treatment

- 1 Give white of eggs beaten with water or skimmed milk
- 2 Dimercaprol (BAL[®]) at once (see page 538)
- 3 Sodium sulfate 30 Gm (1 oz) in water
- 4 Fluids 1000 cc (1 qt) of saline I V at once (may add 1 Gm sodium thiosulfate) and repeat as necessary
- 5 Watch urinary output. Treat oliguria and anuria if it occurs (see page 303)
- 6 Symptomatic and supportive measures as necessary
- 7 In chronic form remove from exposure. may give 10 Gm (15 gr) of sodium thiosulfate in 10 cc (2 1/2 dr) water I V every other day

MORPHINE (AND THE OPIATES) (code No 010 370)

Morphine acts primarily on the C N S causing depression and narcosis. The M L D is 65 mg (1 gr) in susceptible individuals. Manifestations headache nausea excitement depression pin point pupils slow respirations rapid and feeble pulse shock and coma

Treatment

- 1 Nalorphine Hydrochloride U S P (Nallin[®]) an acute antagonist in doses of 5-10 mg I V as an antidote for overdose of morphine and its derivatives meperidine (Demerol[®]) and methadone. If effective in case in pulmonary ventilation is not relieved with the initial dose 5-10 mg may be repeated every 15 minutes until respirations return to normal and patient responds to stimuli
- 2 Maintain adequate respiration by use of artificial respiration or preferably resuscitators with oxygen
- 3 Keep patient awake and warm have him walk if necessary or use ammonia inhalation and strong stimuli
- 4 Lavage stomach well (prevent aspiration) with 1-2000 potassium permanganate at short intervals. Morphine is excreted into the stomach
- 5 Sodium sulfate 30 Gm (1 oz) in water as cathartic
- 6 Atropine sulfate 0.5 mg (1/120 g) subcut if respiratory depression is necessary

MUSHROOMS (code No 010 384)

The Amanita genus of mushrooms accounts for almost all cases of fungus poisoning in the United States. Amanita muscaria poisoning of rapid onset especially ally to atropine if treated promptly recovery most often follows. The deadly type of mushroom poisoning due to Amanita phalloides. A. b. unguis and A. verna has no specific antidote and the prognosis is usually poor. Mushroom poisoning is summarized in the table on page 543

OXALIC ACID (code No 010 3332)

Oxalic acid a component of bleaching powder is a powerful local irritant which precipitates on oxidized calcium. The M L D is 4 Gm (1 dr). Poisoning is manifested by burning in mouth and throat violent abdominal pains bloody vomit and dyspnea tremor oliguria and circulatory collapse

Treatment

- 1 Keep patient quiet in a darkened room avoid stimulation
- 2 At once 0.5 Gm (7 1/2 gr) sodium amytal I.V. slowly in 10-20 cc (2 1/2-5 dr) of water. If not available give drug orally in doses up to 5 times the hypnotic dose. May repeat
- 3 Artificial respiration and oxygen during convulsions
- 4 Lavage gently with 1:2000 potassium permanganate solution before symptoms appear
- 5 Inhalations of ether or chloroform to quiet patient

SNAKE (AND GILA MONSTER) BITES (code No. 010 3814)

The venom of poisonous snakes and lizards may be neurotoxic or hemotoxic. Neurotoxic cause respiratory paralysis hemotoxic cause hemolysis and destruction of endothelial lining of blood vessels. The manifestations are local pain, thirst, profuse perspiration, nausea, vomiting, stimulation followed by depression, local redness, swelling, extravasation of blood and collapse.

Treatment (Avoid opiate and alcohol)

- 1 Keep patient relaxed and quiet
- 2 Apply tourniquet (tight enough to block lymphatic flow but no venous return) above bite releasing for 1-2 minutes every 15-20 minutes
- 3 Cut deep cross incisions at site and apply suction
- 4 Give specific antivenom as soon as possible (Follow printed instructions)
- 5 Plenty of warm fluids
- 6 Biturate sedation as needed
- 7 Institute supportive measures
- 8 Hospitalize and determine blood type as soon as possible give transfusion as necessary. Corticosteroids and antibiotics may be helpful depending on the time of exposure (see pages 423-8)

SPIDER BITES (code No. 010 3815) AND**SCORPION STINGS (code No. 010 3815)****(Black Widow Spider Bite code No. 010 3816)**

The toxin of the less venomous species of spiders and scorpions are usually local pain, redness and swelling. That of the more venomous species causes general muscular pain, convulsions, nausea, vomiting, variable CNS involvement and collapse.

Treatment

- 1 Apply tourniquet cut cross incisions and apply suction
- 2 If absorption has occurred give 10% of 10% calcium gluconate I.V. or I.M. Repeat as necessary
- 3 Give specific antivenom (None available in U.S.A.)
- 4 Keep patient recumbent and quiet
- 5 Hot baths and 20 cc of 10% magnesium sulfate I.V. for relief of pain (see section on page 296)
- 6 Administer dextrose intrathecally supportive measures
- 7 Hot compresses of sodium bicarbonate solution for relief of local pain if systemic involvement
- 8 Corticotropin (ACTH) or other corticosteroids may be of some value in severe cases (see page 423)

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TABLES OF APPROXIMATE EQUIVALENTS

Weight Equivalents		Volume Equivalents	
Apothecary	Metric	Apothecary	Metric
1/320 gr	0.2 mg	1 min (v)	0.05 cc
1/210 gr	0.3 mg	3 min (v)	0.18 cc
1/160 gr	0.4 mg	5 min (v)	0.3 cc
1/120 gr	0.5 mg	8 min (v)	0.5 cc
1/100 gr	0.6 mg	10 min (v)	0.6 cc
1/60 gr	1.0 mg	12 min (v)	0.75 cc
1/30 gr	2.0 mg	15 min (v)	0.9 cc
1/15 gr	4.0 mg	16 min (v)	1.0 cc
1/12 gr	5.4 mg	20 min (v)	1.2 cc
1/10 gr	6.5 mg	30 min (v)	1.8 cc
1/8 gr	8.0 mg	50 min (v)	3.0 cc
1/6 gr	11.0 mg	1 fl dr (3)	3.7 cc
1/4 gr	16.0 mg	65 min (v)	4.0 cc
1/3 gr	22.0 mg	80 min (v)	5.0 cc
3/8 gr	24.0 mg	2 fl dr (3)	7.5 cc
1/2 gr	32.0 mg	2 2/3 fl dr (3)	10.0 cc
3/4 gr	40.0 mg	4 fl dr (3)	15.0 cc
1 gr	65.0 mg	5 1/2 fl dr (3)	20.0 cc
1 1/2 gr	0.1 Gm	8 fl dr (3)	1.0 fl oz
2 gr	0.13 Gm	1 fl o (3)	30.0 cc
3 gr	0.2 Gm	1 2/3 fl oz (3)	50.0 cc
5 gr	0.32 Gm	2 fl o (3)	60.0 cc
1/2 gr	0.5 Gm	3 3/8 fl o (3)	100.0 cc
10 gr	0.85 Gm	4 fl oz (3)	120.0 cc
15 gr	1.0 Gm	8 fl o (3)	240.0 cc
1 dr (3)	4.0 Gm	16 fl o (3)	480.0 cc
1 oz (3)	30.0 Gm	1 pt	480.0 cc

Household Measures	Apothecary	Metric
1 tea spoon	1 fl dr (3)	4 cc
1 table spoon	1/2 fl o (3)	15 cc
1 teacup	4 fl oz (3)	120 cc
1 glass (tumbler)	8 fl oz (3)	240 cc
1 measuring cup	8 fl oz (3)	240 cc
1 pint	16 fl oz (3)	480 cc

CENTIGRADE TO FAHRENHEIT TEMPERATURES

C	F	C	F	C	F
35	95	37.5	99.5	40	104
35.5	95.9	38	100.4	40.5	104.9
36	96.8	38.5	101.3	41	105.8
36.5	97.7	39	102.2	42	107.6
37	98.6	39.5	103.1	43	109.4

METRIC SYSTEM

Weight	1 000 micrograms (γ)	1 milligram (mg)
	1 000 milligrams (mg)	1 gram (Gm)
	1 000 grams (Gm)	1 kilogram (Kg)
Volume	1 000 cubic millimeters	1 milliliter (ml)
		or 1 cubic centimeter (cc)
	1,000 cubic centimeters (cc)	1 liter (L)

IDEAL WEIGHT FOR ADULTS AGES OF 25 AND OVER

(Columns give height in feet and inches and weight in pounds and kilograms)

Height (Feet and Inches)		Ideal Weight for Men					
		Without shoes			With shoes		
Feet	Inches	Lb	Kg	Lb	Kg	Lb	Kg
5	2	114	51.5	124	56.3	131	59.4
5	3	119	54.0	127	57.6	133	60.3
5	4	122	55.8	130	59.0	137	62.1
5	5	126	57.2	134	60.8	141	63.9
5	6	130	58.9	137	62.2	145	65.8
5	7	133	60.3	141	64.0	149	67.6
5	8	136	61.7	145	65.8	153	69.4
5	9	140	63.5	149	67.6	157	71.2
5	10	144	65.3	153	69.4	161	73.0
5	11	148	67.1	157	71.2	165	74.8
6	0	152	68.9	161	73.0	169	76.7
6	1	157	71.2	166	75.3	174	78.9
6	2	163	73.9	171	77.6	179	81.2
6	3	168	76.2	176	79.8	184	83.5

Height (Feet and Inches)		Ideal Weight for Women					
		Without shoes			With shoes		
Feet	Inches	Lb	Kg	Lb	Kg	Lb	Kg
5	0	105	47.6	112	50.8	119	54.0
5	1	107	48.5	114	51.7	121	54.9
5	2	110	49.9	117	53.1	124	56.3
5	3	113	51.3	120	54.4	127	57.6
5	4	116	52.7	124	56.3	131	59.0
5	5	119	54.0	127	57.6	133	60.3
5	6	123	55.8	130	59.0	138	62.1
5	7	126	57.2	134	60.8	141	63.9
5	8	129	58.9	137	62.2	145	65.8
5	9	133	60.3	141	64.0	149	67.6
5	10	137	61.7	145	65.8	153	69.4
5	11	139	63.1	148	67.1	155	70.3
6	0	141	64.0	151	68.5	160	72.6

Feet and inches in parentheses are given in feet and inches. Weights are given in pounds and kilograms. The weight in parentheses is the weight in pounds for a person who is 5 feet 7 inches tall.

AVERAGE HEIGHT AND WEIGHT FOR CHILDREN

Age	Boys				Girls			
	Height	Weight	Height	Weight	Height	Weight	Height	Weight
Years	Feet	Inches	Lb	Kg	Feet	Inches	Lb	Kg
Birth	1	8	45	20	1	8	45	20
1/2	2	2	66	30	2	2	66	30
1	2	5	73	33	2	5	73	33
2	2	8	83	38	2	8	83	38
3	3	0	91	41	3	0	91	41
4	3	3	99	45	3	3	99	45
5	3	6	106	48	3	6	104	47
6	3	9	114	52	3	9	111	50
7	3	11	119	54	3	11	118	53
8	4	2	127	58	4	2	127	58
9	4	4	133	60	4	4	133	60
10	4	6	137	62	4	6	137	62
11	4	8	142	64	4	8	142	64
12	4	10	147	67	4	10	147	67
13	5	0	152	69	5	0	152	69
14	5	2	157	71	5	2	157	71
15	5	4	162	74	5	4	160	73
16	5	6	167	76	5	6	162	74
17	5	8	171	78	5	8	166	76

Weights are given in pounds and kilograms. Heights are given in feet and inches.